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         FEB 20
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NEWS 13 FEB 29
                  WPINDEX/WPIDS/WPIX enhanced with ECLA and current
                  U.S. National Patent Classification
NEWS 14 MAR 31
                  IFICDB, IFIPAT, and IFIUDB enhanced with new custom
                  IPC display formats
NEWS 15 MAR 31
                 CAS REGISTRY enhanced with additional experimental
                  spectra
NEWS 16 MAR 31
                  CA/CAplus and CASREACT patent number format for U.S.
                 applications updated
LPCI now available as a replacement to LDPCI
NEWS 17 MAR 31
NEWS 18 MAR 31
                  EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 19 APR 04
                  STN AnaVist, Version 1, to be discontinued
NEWS 20 APR 15
                  WPIDS, WPINDEX, and WPIX enhanced with new
                  predefined hit display formats
NEWS 21 APR 28
                  EMBASE Controlled Term thesaurus enhanced
NEWS 22 APR 28
                  IMSRESEARCH reloaded with enhancements
NEWS 23 MAY 30
                  INPAFAMDB now available on STN for patent family
                  searching
                  DGENE, PCTGEN, and USGENE enhanced with new homology
NEWS 24 MAY 30
                  sequence search option
NEWS 25 JUN 06 EPFULL enhanced with 260,000 English abstracts
NEWS 26 JUN 06 KOREAPAT updated with 41,000 documents
NEWS EXPRESS FEBRUARY 08 CURRENT WINDOWS VERSION IS V8.3.
              AND CURRENT DISCOVER FILE IS DATED 20 FEBRUARY 2008
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89.6% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

24 ANSWERS

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FULL FILE PROJECTIONS: ONLINE **COMPLETE**
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24 SEA SSS SAM L1

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FULL SCREEN SEARCH COMPLETED - 44523 TO ITE 44523 TO ITERATE

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=> file caplus

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                  (PESTIVIRUS OR PESTIVIRUSES)
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          14187 HCV
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                  (HEPATITIS OR HEPATITISES)
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=> s 14 and (flavivirus or pestivirus or flaviviridae or how or hepatitis c)
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           2025 FLAVIVIRUS
                  (FLAVIVIRUS OR FLAVIVIRUSES)
            501 PESTIVIRUS
            266 PESTIVIRUSES
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                  (PESTIVIRUS OR PESTIVIRUSES)
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                  (HEPATITIS OR HEPATITISES)
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                  (HEPATITIS (W) C)
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58 L4 AND (FLAVIVIRUS OR PESTIVIRUS OR FLAVIVIRIDAE OR HCV OR HEPAT

ITIS C)

=> d bib abs hitstr 40-58 16

ANSWER 40 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN AN 2006:103884 CAPLUS

144:171198 DN

Preparation of alkyl-substituted 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs via condensation of the lactone to nucleosides as potential antiviral agents
Wang, Peiyuan; Stec, Wojciech; Clark, Jeremy; Chun, Byoung-Kwon; Shi,

IN

Junxing; Du, Jinfa Pharmasset, Inc., USA PA

PCT Int. Appl., 34 pp. CODEN: PIXXD2

DТ Patent

LA English FAN.CNT 1

	PATENT NO.									APPLICATION NO.						DATE					
PI	WO	2006012440 2006012440				A2					WO	2005	-US25	916		2	20050721 \$E, CA, CH, CH, CA, CH, CH, CA, CH, CH, CH, CH, CH, CH, CH, CH, CH, CH				
		W:	CN, GE, LC, NG,	CO, GH, LK, NI,	CR, GM, LR, NO,	CU, HR, LS, NZ,	CZ, HU, LT, OM,	DE, ID, LU, PG,	DK, IL, LV, PH,	DM, IN, MA, PL,	D2 IS MI P1	, EG	KE, KE, MK,	EG, KG, MN, SC,	ES, KM, MW, SD,	FI, KP, MX, SE,	GB, KR, MZ, SG,	GD, KZ, NA, SK,			
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											CA 2005-2574651										
	EP	1773								EP 2005-775359											
		R:																			
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	BR	2005	0121	04		A		2008	0311		BR	2005	-1210	14		2	0050	721			
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	KB	2007	1143	44		A		2007	1203		KB	200	7-7039	180		2					
PRAT	US	2007 2004	-589	866P		P		2004	0721												
	US	2004	-608	320P		P		2004	0909												
	US	2005	-185	988		A1		2005	0721												
	WO	2005	-US2	5916		W		2005	0721												

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB A process for preparing of 2-deoxy-2-fluoro-2-methyl-D-ribonolactones, I, wherein R1 and R2 can independently be H, CH3, acetyl, benzoyl, pivaloyl, 4-nitrobenzoyl, 3-nitrobenzoyl, 2-nitrobenzoyl, 4-chlorobenzoyl, 3-chlorobenzoyl, 2-chlorobenzoyl, 4-methylbenzoyl, 3-methylbenzoyl, 2-methylbenzoyl, 4-phenylbenzoyl, benzyl, 4-methoxybenzyl, trityl, trialkylsilyl, t-butyl-dialkylsilyl, t-butyldiphenylsilyl, TIPDS, THP, MOM, or MEM are prepared and used in the condensation to 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs. Thus, 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs II and III, wherein X is a halogen; Y is N or CH; Z is a halogen hydroxyl, ether, thiol, thioether, (un) substituted amine or alkyl; Rl' is alkyl, vinyl, ethynyl; R2' and R3' can be same or different H, alkyl, arylalkyl, acyl, cyclic acetal such as 2',3'-O-isopropylidene or 2',3-O-benzylidene, or 2',3'-cyclic carbonate; R4, R5, and R6 are independently H, halogen, hydroxyl, ether, thiol, thioether, N3,

os

GT

MARPAT 144:171198

(un) substituted amine, (un) substituted amido, alkyl, halogenated alkyl, alkenyl, halogenated alkenyl, alkynyl, hydroxy alkyl, alkoxy are prepared and are potential anti-HCV agents.

Specifically, IV was prepared (no yield, claimed) via condensation, alkylation and stereoselective fluorimation reactions and can exhibit

potential use as an anti-HCV agent. T 874638-81-0P

RI: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of alky-substituted Z-deoxy-Z-liuoro-D-inofuranosyl pyrimidine and purine nucleoside analogs via condensation of the lactone to nucleosides)

RN 874638-81-0 CAPLUS

CN Benzamide, N-[1-[5-0-benzoyl-2-C-methyl-3-0-(methylsulfonyl)-\(\beta\)-Darabinofuranosyl]-1,2-dihydro-2-oxo-4-pyrimidinyl]- (CA INDEX NAME)

Absolute stereochemistry.

- L6 ANSWER 41 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2005:1151389 CAPLUS
- DN 145:271979
- TI NM 283, an efficient prodrug of the potent anti-HCV agent
 - 2'-C-methylcytidine
- AU Pierra, C.; Benzaria, S.; Amador, A.; Moussa, A.; Mathieu, S.; Storer, R.; Gosselin, G.
- CS Laboratoire Cooperatif Idenix, CNRS, Universite Montpellier II.
- Montpellier, 5, Fr.
- SO Nucleosides, Nucleotides & Nucleic Acids (2005), 24(5-7), 767-770 CODEN: NNNAFY; ISSN: 1525-7770
- PB Taylor & Francis, Inc. DT Journal
- DT Journal LA English
- OS CASREACT 145:271979
- AB In order to improve the oral bloavailability of 2"-C-methyloytidine, a potent anti-ERV agent, the corresponding 3"-C-I-valinyl estimates and its physicochem. properties, Mr 283 has seen synthesized. Based on its ease of synthesis and its physicochem. properties, Mr 283 has emerged as a promising antiviral drug for treatment of chronic BCV infection. 22643-36-99 31448-34-19 640753-79 21469-34-19 640754-79
- IT 23643-36-9P 314 642075-42-1P
 - RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 - (preparation of NM 283 as efficient prodrug of potent anti-HCV
- agent 2'-C-methylcytidine)
- RN 23643-36-9 CAPLUS
 - N 2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-benzoyl-2-C-methyl-β-D-ribofuranosyl)- (9CI) (CA INDEX NAME)

RN 31448-54-1 CAPLUS CN Uridine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

Absolute stereochemistry.

642075-42-1 CAPLUS

Cytidine, N-[(dimethylamino)methylene]-2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

20724-73-6P RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent);

10/609.298

USES (Uses)

(preparation of NM 283 as efficient prodrug of potent anti-HCV agent 2'-C-methylcytidine) 20724-73-6 CAPLUS

Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

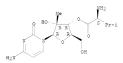
640725-71-9P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); Nai raw (properties); SFN (Synthetic preparation); THU (Therapeutic use BIOL (Biological study); PREP (Preparation); USES (Uses) (producy; preparation of NN 283 as efficient producy of potent anti-HCV agent 2"-C-methylcytidine)

640725-71-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



■ 2 HC1

RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 42 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:684531 CAPLUS

143:431740

Emerging drugs for chronic hepatitis C

Bhopale, Girish Mahadeorao; Nanda, Rabindra Kumar Research and Development Division, Hindustan Antibiotics Limited, Pimpri, CS Pune, 411018, India

Hepatology Research (2005), 32(3), 146-153 CODEN: HPRSFM: ISSN: 1386-6346 so

Elsevier B.V.

PB Journal; General Review

LA English

A review. Hepatitis C virus (HCV) is a

major cause of chronic hepatitis, liver cirrhosis and hepatocellular carcinoma worldwide. A combination therapy comprising pegylated interferon and ribavirin currently represents the most effective therapy for chronic HCV infection. The limitations of this current therapy mainly its efficacy and significant side effects have prompted the development of new drugs. Few categories of therapeutic agents appear promising for future therapy, e.g. novel interferons, ribavirin analogs, antisense oligonucleotides, short interfering RNAs, ribozymes, enzyme inhibitors, immunomodulatory agents, antifibrotic agents, therapeutic vaccines and antibodies. Few drugs belong to afore-mentioned categories have already reached the different clin. phases of development. The

10/609.298

present article highlights the status of current available therapies and emerging drugs for the treatment of hepatitis C.

640725-71-9, NM 283 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(NM283 proved promising therapeutic effect in treating chronic hepatitis C patient) 640725-71-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HCl

THERE ARE 64 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 43 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:648160 CAPLUS 143:248607 DN

Design, Synthesis, and Antiviral Activity of 2'-Deoxy-2'-fluoro-2'-Cmethyl-cytidine, a Potent Inhibitor of Hepatitis C Virus Replication

Clark, Jeremy L.; Hollecker, Laurent; Mason, J. Christian; Stuyver, Lieven J.; Tharnish, Phillip M.; Lostia, Stefanla; McBrayer, Tamara R.; Schimazi, Raymond F.; Watanabe, Kyoichi A.; Otto, Michael J.; Furman, Phillip A.; AU Stec, Wojciech J.; Patterson, Steven E.; Pankiewicz, Krzysztof W. Pharmasset, Inc., Princeton, NJ, 08540, USA Journal of Medicinal Chemistry (2005), 48(17), 5504-5508 00

CODEN: JMCMAR; ISSN: 0022-2623

American Chemical Society

Journal īΑ English

os CASREACT 143:248607

The pyrimidine nucleoside- β-D-2'-deoxy-2'-fluoro-2'-C-methylcytidine

(I) was designed as a hepatitis C virus RNA-dependent

RNA polymerase (HCV RdRp) inhibitor. The title compound was obtained by a DAST fluorination of N4-benzoyl-1-(2-methyl-3,5-di-0-benzoyl-B-D-arabinofuranosv1)cvtosine to provide N4-benzov1-1-(2-fluoro-2 methyl-3,5-di-O-benzoyl-β-D-ribofuranosyl)cytosine. The protected 2'-C-methylcytidine was obtained as a byproduct from the DAST fluorination

and allowed for the preparation of two biol. active compds. from a common precursor. Compound I and 2 -C-methylcytidine were assayed in a sub-genomic HCV replicon assay system and found to be potent and selective inhibitors of HCV replication. Compd.I shows increased

inhibitory activity in the HCV replicon assay compared to 2'-C-methylcytidine and low cellular toxicity.

20724-73-6

RL: PAC (Pharmacological activity); BIOL (Biological study) (design, synthesis via fluorination, and antiviral activity of 2'-deoxy-2'-fluoro-2'-C-methyl-cytidine, a potent inhibitor of Hepatitis C virus replication)

20724-73-6 CAPLUS Cytidine, 2'-C-methyl- (CA INDEX NAME) DM

CN

817204-35-6P 863329-62-8P 863329-64-0P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT

(Reactant or reagent) (design, synthesis via fluorination, and antiviral activity of 2'-deoxy-2'-fluoro-2'-C-methyl-cytidine, a potent inhibitor of Hepatitis C virus replication) 817204-35-6 CAPUS

Senzamide, N-[1,2-dihydro-1-(2-C-methyl- β -D-arabinofuranosyl)-2-oxo-4-pyrimidinyl]- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN

863329-62-8 CAPLUS Benzamide, N-[1-(3,5-di-O-benzoyl-2-C-methyl- β -D-arabinofuranosyl)-1,2-dihydro-2-oxo-4-pyrimidinyl]- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

863329-64-0 CAPLUS

Cytidine, N-benzoyl-2'-C-methyl-, 3',5'-dibenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 19 ALL CITATIONS AVAILABLE IN THE RE FORMAT

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ANSWER 44 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
     2005:34765 CAPLUS
AN
     Preparation of modified fluorinated (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl
     nucleoside analogs as antiviral agents
     Clark, Jeremy
Pharmasset, Ltd., Barbados
IN
     PCT Int. Appl., 228 pp.
     CODEN: PIXXD2
DT
     Patent
A.T
     English
FAN.CNT 1
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              GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
              LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
              NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
              IJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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AB The disclosed invention provides nucleoside analogs I, wherein B is purine and pyrindidne nucleobase; X is O, S, CHZ, Os, NH, N-alkyl, CHM, C(M); Z W is F, Cl, Br, lodo; Rl is H, phosphate, H-phosphonate, acyl, Fh, alkyl, orbowyglkylamino, sulfonate ester, peptide, anino acid, sugar reside; R2 and R

marrow cells are reported. (2'R)-2'-deoxy-2'-fluoro-2'-C-methylcytidine shows activity against Rhinovirus, West Nile virus, Yellow Fever virus, and Dengue virus. Cytotoxicity and effect of nucleoside analogs on human bone marrow cells are reported.

20724-73-6 374750-28-4

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (preparation of modified fluorinated (2'R)-2'-deoxy-2'-fluoro-2'-C-Me nucleoside analogs as antiviral agents)

20724-73-6 CAPLUS Cytidine, 2'-C-methyl- (CA INDEX NAME) CN

Absolute stereochemistry.

374750-28-4 CAPLUS

Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

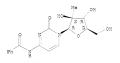
817204-35-6P 817204-36-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of modified fluorinated (2'R)-2'-deoxy-2'-fluoro-2'-C-Me nucleoside analogs as antiviral agents) 817204-35-6 CAPLUS

Benzamide, N-[1,2-dihydro-1-(2-C-methyl- β -D-arabinofuranosyl)-2-oxo-4pyrimidinyl] - (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



817204-36-7 CAPLUS DAT

Benzamide, N-[1,2-dihydro-1-[2-C-methyl-3,5-bis-0-(trifluoroacetyl)- β -D-arabinofuranosyl]-2-oxo-4-pyrimidinyl]- (9CI) (CA INDEX NAME)

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Ph K
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ANSWER 45 OF 58 CAPLUS COPYRIGHT 2008 ACS on SIN
      2004:817630 CAPLUS
AN
DN
      141:307495
      Use of nucleoside compounds and PALA for the treatment of
      flaviviridae infections
     Stuyver, Lieven J.
     Pharmasset Ltd., Barbados
PCT Int. Appl., 120 pp.
PA
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT
      PATENT NO.
                            KIND
                                    DATE
                                                  APPLICATION NO.
                                                                             DATE
      WO 2004084796
                                     20041007
                                                                             20040329
PI
                              A2
                                                  WO 2004-IB1429
      WO 2004084796
                              A3
                                     20060406
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
               CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
               GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
               LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
               NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
               TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
          RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
               BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
               ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, FL, FT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
               TD, TG
     AU 2004224575
                              A1
                                     20041007
                                                   AU 2004-224575
     CA 2529311
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                                                                             20040329
                              A1
                                                  US 2004-812448
EP 2004-724085
      US 20050049204
                              Al
                                                                              20040329
     EP 1626692
                              A2
                                     20060222
                                                                             20040329
          R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK
                                     20060704
      BR 2004008846
                                                  BR 2004-8846
JP 2006-506588
CN 2004-80011746
                                                                             20040329
                              A
      JP 2006524227
                                                                              20040329
      CN 1980678
                                     20070613
                              А
     MX 2005PA10419
                              Δ
                                     20060531
                                                  MX 2005-PA10419
                                                                             20050928
PRAI US 2003-458635P
                              P
     WO 2004-IB1429
                              1.2
                                     20040329
     MARPAT 141:307495
     The invention discloses a composition for and a method of treating
      Flaviviridae infections, e.g. bovine viral diarrhea virus, dengue
     Virus, West Nile virus, and hepatitis C virus, as well as abnormal cellular proliferation, in a host, including animals, and especially
     humans, using a nucleoside compound (Markush included) or
      N-(phosphonoacetyl)-L-aspartate (PALA), or a pharmaceutically acceptable
      salt or prodrug thereof.
      20724-73-6
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (nucleoside compds, and PALA for treatment of flaviviridae
         infections)
      20724-73-6 CAPLUS
     Cytidine, 2'-C-methyl- (CA INDEX NAME)
Absolute stereochemistry.
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2004:780543 CAPLUS
AN
DN
     141:296247
     Preparation of cytidine nucleoside analogs as antiviral agents
     Girardet, Jean-Luc; Koh, Yung-Hyo; An, Haoyun; Hong, Zhi
Ribapharm Inc., USA
IN
PA
so
     PCT Int. Appl., 59 pp.
CODEN: PIXXD2
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                           KIND
                                  DATE
                                                APPLICATION NO.
                                                                         DATE
     WO 2004080466
                            A1
                                   20040923
                                                WO 2003-US6992
                                                                         20030307
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              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
              PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN,
                                                                       TR, TT,
              UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
              FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
              BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
225705 Al 20040930 AU 2003-225705 20030307
     AU 2003225705
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ANSWER 46 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

Α

PRAI WO 2003-US6992

MARPAT 141:296247

os

GT

AB Cytidine analogs I, wherein -KeY- is -NeW, -CHeNT, -NeCZ- or -CHeCZ-, wherein Z is B, halogen, or alkyl, and wherein M is a sugar or sugar analog; wherein the compound has a D-configuration or an I-configuration; with the proviso that where M is a substituted sugar with a ribofurance ring having a heteroatom and substituents R1 and R2 on the C3'-atom, R3 and R4 on the C3'-atom, R3 and R4 on the C3'-atom, and R5 on the C5'-atom, R1-R4 together are not independently H, OH, F, NR2, N3, O-hydrocarbyl, or a reporter molety, when the heteroatom is O, S, Se, SO, N-alkyl, or CR2, and when R5 is OH, SH, NR2, monophosphate, diphosphate, triphosphate, thin NR2, monophosphate, of the composition of the M3 of the C5' of the C5

10/609,298

nucleoside analog II was prepared and tested as antiviral agent. The virus is an HCV virus, an HIV virus, an RSV virus, an influenza virus, or a an HBV virus. 23643-36-9

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of cytidine nucleoside analogs as antiviral agents) 23643-36-9 CAPJUS

2.4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-0-benzoyl-2-C-methyl- β -D-ribofuranosyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

760965-52-4P 760965-53-5P 760965-55-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of cytidine nucleoside analogs as antiviral agents) 760965-52-4 CAPLUS Uridine, 5-bromo-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

760965-53-5 CAPLUS Cytidine, 5-bromo-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

760965-55-7 CAPLUS

Cytidine, 5-amino-2'-C-methyl- (9CI) (CA INDEX NAME)

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNI 9 ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 47 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

2004:652668 CAPLUS

DN 141:167739

Diribonucleotides as specific viral RNA-polymerase inhibitors for the treatment or prevention of viral infections Wu, Jim Zhen; An, Haoyun; Hong, Zhi

IN PA USA

U.S. Pat. Appl. Publ., 12 pp.

CODEN: USXXCO Patent

English LA

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
PI PRAI OS	US 20040158054 US 2003-360218 MARPAT 141:167739	Al	20040812 20030207	US 2003-360218	2003020		

- etc.) comprising a first and second nucleoside. The dinucleotide inhibits viral RNA polymerase and at least one of the nucleosides exhibits _____ polymerase and at least one of the nucleosid antiviral activity when cleaved from the dinucleotide. 735268-87-87
- RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
 - (diribonucleotides as specific viral RNA-polymerase inhibitors for treatment or prevention of viral infections) 735268-87-8 CAPLUS

Cytidine, 2'-0-methylguanylyl-(3'-5')-2'-C-methyl- (9CI) (CA INDEX

Absolute stereochemistry.

- 735268-88-9 RL: RCT (Reactant); RACT (Reactant or reagent) (diribonucleotides as specific viral RNA-polymerase inhibitors for treatment or prevention of viral infections) 735268-89-9 CAPLUS
- RN Cytidine, N-benzoyl-5'-0-[bis(4-methoxyphenyl)phenylmethyl]-P(0)-(2-cyanoethyl)-P-doxo-2'-0-methylguanylyl-(3'-5')-N-benzoyl-2'-C-methyl-gyanylyl-(3'-5')-N-benzoyl-2'-C-methyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

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ANSWER 48 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
1.6
AN
     2004:566635 CAPLUS
DN
      141:89323
     Process for the production of 3'-nucleoside prodrugs
TN
      Storer, Richard; Moussa, Adel; Mathieu, Steven; Qu, Lin
PA
      Idenix Cayman Limited, Cayman I.
SO
     PCT Int. Appl., 57 pp.
CODEN: PIXXD2
DT
      Patent
LA
      English
FAN.CNT 1
      PATENT NO.
                            KIND
                                    DATE
                                                  APPLICATION NO.
                                                                             DATE
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DT
    WO 2004058792
                             A1
                                     20040715
                                                  WO 2003-US41603
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               GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
               LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
               NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
          TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
               BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
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     CA 2511616
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AU 2003-300434
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      EP 1575971
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                                                                             20031223
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      CN 1751058
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      JP 2006514038
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      NZ 540913
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      ZA 2005005040
                                     20060426
                                                  ZA 2005-5040
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      NO 2005003557
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PRAI US 2002-436150P
     WO 2003-US41603
                             W
os
     CASREACT 141:89323; MARPAT 141:89323
GΙ
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10/609.298

- AB Provided is a single-step process for the regionelective 3'-acylation of a ribofurnacyl' 2'- or 3'-branched nucleosides 1', wherein B is nucleobase. These compds are useful as antiviral agents, and in particular, can be used to treat Flavivirdse infections in a host in need thereof (no data). Thus, 9-(2'-C-methyl-3'-O-valincyl-B-D-ribofurnacyl)-6-N-methyladenine dhydrocoloride was prepared via regionelective esterification of 9-(2'-C-methyl-B-D-ribofurnacyl)-6-N-methyladenine with N-(ter-boutscyparhoppyl)-1-valine.
 - 640725-70-8P
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
 - (process for production of nucleoside prodrugs via regioselective esterification)
 - RN 640725-70-8 CAPLUS
 - I L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

- IT 20724-73-6
 - RL: RCT (Reactant); RACT (Reactant or reagent)
 (process for production of nucleoside prodrugs via regioselective
- esterification)
 RN 20724-73-6 CAPIUS
- RN 20724-73-6 CAPLUS CN Cytidine, 2'-C-methyl- (CA INDEX NAME)

- 16 ANSWER 49 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:453348 CAPLUS
- DN 141:17578
- II Treatment of Flaviviridae infection with 2'-branched nucleosides
- and another mutation-inducing drug such as interferon
- IN Sommadossi, Jean-Pierre; La Colla, Paolo; Standring, David; Bichko, Vadim; Qu, Lin
- PA Idenix (Cayman) Limited, Cayman I.; Universita Degli Studi Di Cagliari SO PCI Int. Appl., 166 pp.
- CODEN: PIXXD2 DI Patent
- DT Patent LA English
- FAN.CNT 1
 PATENT NO. KIND DATE APPLICATION NO. DATE

 PI NO 2004046331 A2 20040603 MO 2003-US36714 20031117
 NO 2004046331 A3 20063002
 - WC 2004046331 A3 20060302 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

AB

CN

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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE,
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              LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
              OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
                                                                                TM.
              TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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              TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD,
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     CA 2506129
                            A1
                                                                         20031117
     AU 2003298658
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                                                AU 2003-298658
     US 20050031588
                            Al
                                   20050210
                                               US 2003-715729
EP 2003-796412
     EP 1576138
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     BR 2003016363
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                                   20051004
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     JP 2006519753
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     CN 1849142
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                                   20061018
                                                CN 2003-80108747
     MX 2005PA05192
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     NO 2005002920
                                               NO 2005-2920
                                   20050815
                                                                         20050615
                            Α
PRAI US 2002-426675P
                            P
     WO 2003-US36714
     MARPAT 141:17578
     The present invention discloses a method for the treatment of
     Flaviviridae infection that includes the administration of a
     2'-branched nucleoside, or a pharmaceutically acceptable prodrug and/or
     salt thereof, to a human in need of therapy in combination or alternation
     with a drug that directly or indirectly induces a mutation in the viral
     genome at a location other than a mutation of a nucleotide that results in
     a change from serine to a different amino acid in the highly conserved
     consensus sequence, XRX<u>S</u>GXXXT, of domain B of the RNA polymerase
     region, or is associated with such a mutation. The invention also includes a
     method to detect a mutant strain of Flaviviridae and a method
     for its treatment. Thus, in bovine viral diarrhea virus (BVDV)-infected
     MDBK cells treated with \beta-D-2'-methylcytidine, viruses resistant to
     the nucleoside appeared. The drug resistance was associated with a mutation
     in the NS5B gene which resulted in an S405T substitution in the encoded
     RNA-dependent RNA polymerase. These mutant viruses were sensitive to Intron A (interferon \alpha-2b). Intron A and \beta-D-2'-methylcytidine
     exhibited synergistic inhibitory activity on BVDV growth in MDBK cells.
     20724-73-6
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (treatment of Flaviviridae infection with 2'-branched
        nucleosides and another mutation-inducing drug such as interferon)
    20724-73-6 CAPLUS
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Absolute stereochemistry.

640281-90-9

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (treatment of Flaviviridae infection with 2'-branched nucleosides and another mutation-inducing drug such as interferon) 640281-90-9 CAPLUS CNI L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

Cytidine, 2'-C-methyl- (CA INDEX NAME)

- ANSWER 50 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN 2004:20697 CAPLUS 140:87662
- AN DN
- 2'- and 3'-nucleoside prodrugs for treating Flaviviridae
- infections
- IN Sommadossi, Jean-pierre; La Colla, Paolo; Storer, Richard; Gosselin, Gilles
- Genius (Ayman) Limited, Cayman I.; Centre National de la Recherche Scientifique; Universita Degli Studi di Cagliari PCI Int. Appl., 2498 pp.
 CODEN: PIXXD2 PA
- so
- Patent
- English

	PA:	TENT		KIN		DATE			APP:	LICAT	ION	NO.		D.	ATE					
PI		2004				A2		20040108		WO 2003-IB3901						20030627				
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												, MW,								
												, SG,				TJ,	TM,	TN,		
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	CN	1011	7299	3		A		2008				2007-				2	0030	627		
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				TD,	TG															
	EP	1656				A2						2004-					0040			
		R:										, IT,								
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		2007				A1		2007				2004-					0041 0041			
				400		A1 A1		2007	0201		U.S.	2004- 2004-	24/0							
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HR

PRA:

		20070032407 7192936	Al B2	20070208	US	2004-5473	20041206
		20070037735	Al	20070215	IIC	2004-5442	20041206
		20070042939	A1	20070222		2004-5445	20041206
		20070042991	Al	20070222		2004-5447	20041206
		7365057	B2	20080429	0.0	2004-3441	20041200
		20070042940	A1	20070222	TIS	2004-5467	20041206
		20070042990	Al	20070222		2004-5471	20041206
		20070060503	A1	20070315		2004-5440	20041206
		20070060498	Al	20070315		2004-5444	20041206
	US	20070060504	A1	20070315	US	2004-5446	20041206
	US	20070060541	Al	20070315	US	2004-5466	20041206
		20070060505	A1	20070315	US	2004-5472	20041206
	MX	2004PA12779	A	20050819	MX	2004-PA12779	20041216
	NO	2005000466	A	20050323	NO	2005-466	20050127
		2005DN00341	A	20070202	IN	2005-DN341	20050128
		20070275883	A1	20071129	US	2006-516928	20060906
ıΙ		2002-392350P	P	20020628			
		2002-392351P	P	20020628			
		2003-466194P	P	20030428			
		2003-470949P	P	20030514			
		2003-820501	A3	20030627			
		2003-820701	A3	20030627			
		2003-607909	A1	20030627			
		2003-608907	A1	20030627			
		2003-609298	A1	20030627			
		2003-IB3901	W	20030627			
		2004-US15395	W	20040514			
	MAI	RPAT 140:87662					

AB 2' And 3'-Prodrugs of 1'-, 2'-, 3'-, or 4'-branched β-D or β-L nucleosides, or their pharmaceutically acceptable salts and derivs., are described which are useful in the prevention and treatment of Flaviviridae infections and other related conditions. These modified nucleosides provide superior results against flaviviruses and pestiviruses, including hepatitis virus and viruses generally that replicate through an RNA-dependent RNA reverse transcriptase. Compds., compns., methods and uses are provided for the treatment of Flaviviridae infection, including HCV infection, that include the administration of an effective amount of the prodrugs of the invention, or their pharmaceutically acceptable salts or derivs. These drugs may optionally be administered in combination or alternation with further antiviral agents to prevent or treat Flaviviridae infections and other related conditions. Preparation of compds. of the invention is included.

20724-73-GF RE: ADV (Adverse effect, including toxicity); BSU (Biological study, unclassified); DNA (Drug mechanism of action); PAC (Pharmacological activity); PAT (Pharmacokinetics); RCT (Reactant); SPN (Synthetic preparation); TNY (Therapeutic use); BTOL (Biological study); PREP (Incleaning through of the through Flavivirides infections)

(nucleoside prodrugs for treating Flaviviridae infections)
20724-73-6 CAPLUS

Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

IT 125911-78-6 386213-38-3

RL: BSU (Biological study, unclassified); BIOL (Biological study) (nucleoside prodrugs for treating Flaviviridae infections)

RN 125911-78-6 CAPLUS

CN 5'-Uridylic acid, 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

386213-38-3 CAPLUS 5'-Cytidylic acid, 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

125911-76-4 150993-73-0 640725-72-0

RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); BIOL (Biological study) (nucleoside prodrugs for treating Flaviviridae infections)

125911-76-4 CAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

150993-73-0 CAPLUS

Uridine 5'-(trihydrogen diphosphate), 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 640725-72-0 CAPLUS CN Cytidihe 5'-(trihydrogen diphosphate), 2'-C-methyl- (9CI) (CA INDEX NAME)

- 374750-28-4
 RI: BEU (Blological study, unclassified); PAC (Pharmacological activity);
 FRI (Pharmacokinetics); BIOL (Blological study)
 (nucleoside prodrugs for treating Flaviviridae infections)
- 374750-28-4 CAPLUS
- Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- 640725-71-9P
 - RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (nucleoside prodrugs for treating Flaviviridae infections)

- 640725-71-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



- 31448-54-1 188413-99-2
- RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nucleoside prodrugs for treating Flaviviridae infections)
- 31448-54-1 CAPLUS Uridine, 2'-C-methyl- (CA INDEX NAME) RN
- CN
- Absolute stereochemistry. Rotation (+).

RN

188413-99-2 CAPLUS Cytidine, 2'-C-ethynyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

640725-69-5P 640725-70-8P

- RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
- (nucleoside prodrugs for treating Flaviviridae infections)
 640725-69-5 CAPLUS
 Cytidine, 2'-C-methyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

640725-70-8 CAPLUS L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

McIntosh

IT 622381-09-3

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological atudy); USES (Uses) (nucleoside prodrugs for treating Flaviviridae infections,

and use with other agents)

622381-09-3 CAPLUS

CN Cytidine, N-cyclopropyl-2'-C-methyl- (9CI) (CA INDEX NAME)

- 16 ANSWER 51 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:20696 CAPLUS DN 140:77365
- TI Preparation of modified 2'- and 3'-nucleoside prodrugs for treating Flaviviridae infections
- Sommadossi, Jean-pierre; La Colla, Poalo; Storer, Richard; Gosselin, IN
- Clemix (Cayman) Limited, Cayman I.; Universita degli studi di Cagliari; Centre National de la Recherche Scientifique PCT Int. Appl., 201 pp. CODEN: FIXO. PA so
- DT Patent
- LA English

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EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20060517 ED 1656093 20 FD 2004_776022 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HF US 20070027065 US 20070027104 US 2004-5468 20041206 A1 US 2004-5469 20041206 2.1 US 20070027066 US 2004-5470 A1 20041206 US 20070032449 US 2004-5441 20041206 A1 US 20070032407 A1 20070208 US 2004-5473 20041206 US 7192936 B2 A1 US 2004-5442 20041206 US 20070042939 US 2004-5445 20041206 A1 US 20070042991 US 2004-5447 20041206 2.1 US 7365057 B2 20080429 US 20070042940 A1 US 2004-5467 20041206 US 20070042990 A1 US 2004-5471 20041206 US 20070060503 21 20070315 US 2004-5440 20041206 US 2004-5444 US 2004-5446 20041206 US 20070060498 A1 US 20070060504 A1 20041206 US 20070060541 US 2004-5466 20041206 A1 US 2004-5472 MX 2004-PA12709 US 20070060505 A1 20041206 MX 2004PA12709 Α 20050930 20041215 NO 2005000465 Α NO 2005-465 US 2006-516928 20050127 US 20070275883 Al 20071129 20060906 P PRAI US 2002-392350P 20020628 US 2002-392351P P US 2003-466194P P 20030428 US 2003-470949P P 20030514 CN 2003-820501 A3 20030627 CN 2003-820701 A3 tts 2003-607909 A1 US 2003-608907 A1 20030627 US 2003-609298 20030627 A1 WO 2003-IB3246 7.7 WO 2004-US15395 w 20040514 MARPAT 140:77365

GT

2' And/or 3' prodrugs of 1', 2', 3' or 4'-branched-nucleosides I, wherein R1-R3 are independently H, phosphate, alkyl, acyl, C0-alkyl, C0-aryl, CO-alkoxyalkyl, CO-aryloxyalkyl, CO-substituted aryl, sulfonate ester, benzyl, wherein the Ph group is optionally substituted with one or more substituents, alkylsulfonyl, arylsulfonyl, aralkylsulfonyl, lipid, amino acid, carbohydrate, peptide, cholesterol; Y1 is hydrogen, bromo, chloro, fluoro, iodo, CN, OH, OR4, NH2, NHR4, NR4R5, SH or SR4; X1 and X2 are Tidoro, 2000, U.N., U.N., U.N., U.N., NARAN, NARAS, SA OF SAY, AI and AL are independently alkyl, CHR, CF3, CF3, 73, 2-Br-Et, CH2P, CH2C1, CH2CF3, CF2CF3, CY2CY3, CH2CH, alkenyl, alkynyl, COOH, COORA, COO-alkyl, COO-aryl, COO-o-alkoyalkyl, CONH2, CONH34, CON (R4) Z, halo, CN, N3, OH, ORA, NH2, NHR4, NH4R5, SH or SR5; Y Ls independently H, halo; and each R4 and R5 is independently hydrogen, acyl, alkyl, lower alkyl, alkenyl, alkynyl or cycloalkyl, and their pharmaceutically acceptable salts and derivs. are described. These prodrugs are useful in the prevention and treatment of Flaviviridae infections, including BCV infection, and other related conditions. Compds. and compns. of the prodrugs of the

present invention are described. Methods and uses are also provided that include the administration of an effective amount of the prodrugs of the present invention, or their pharmaceutically acceptable salts or derivs. These drugs may optionally be administered in combination or alteration with further anti-viral agents to prevent or treat Flaviviridae infections and other related conditions. Thus, antiviral activity of β-D-2'-C-methyl-7-methyl-6-phenyl-3,3a,5,8a-tetrahydro-1,3,4,5,7apenta-aza-s-indacen-8-one is reported.

640281-90-9P

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of modified and nucleoside prodrugs for treating flaviviridae infections)

640281-90-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

20724-73-6

RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of modified and nucleoside prodrugs for treating flaviviridae infections)

20724-73-6 CAPLUS Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

ANSWER 52 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

KIND DATE

- AN 2004:20443 CAPLUS
- DN 140:70984
- 2'-C-methyl-3'-O-L-valine ester ribofuranosyl cytidine for treatment of flaviviridae infections
- Sommadossi, Jean-Pierre; La Colla, Paolo Idenix (Cayman) Limited, Cayman I.; Universita Degli Studi di Cagliari PCT Int. Appl., 110 pp. PA
- so CODEN: PIXXD2
- Patent
- English FAN.CNT 4

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	WO	2004002422				A3 20050407												
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APPLICATION NO.

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              NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
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PRAI US 2002-392351P
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     WO 2004-US15395
                                   20040514
                           W
      IN 2005-DN344
                            A3
                                   20050128
     MARPAT 140:70984
     The 3'-L-valine ester of β-D-2'-C-methyl-ribofuranosyl cytidine
     provides superior results against flaviviruses and
     pestiviruses, including hepatitis C virus.
     Based on this discovery, compds., compns., methods and uses are provided for the treatment of flaviviridae, including HCV, that
     include the administration of an effective amount of val-mCvd or its salt,
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ester, prodrug or derivative, optionally in a pharmaceutically acceptable
      carrier. In an alternative embodiment, val-mcyd is used to treat any virus that replicates through an RNA-dependent RNA polymerase. Several
      examples are provided of the pharmacol., mechanism of action, metabolism, side
      effects, and clin. efficacy of the title compound
      640281-90-9D, salts 642075-50-1 642075-51-2
      642075-52-3 642075-53-4 642075-54-5
642075-55-6 642075-56-7 642075-57-8
642075-58-9 642075-59-0 642075-60-3
      642075-61-4 642075-62-5 642075-63-6
      642075-64-7 642075-65-8 642075-66-9
642075-67-0 642075-68-1 642075-69-2
642075-70-5 642075-71-6 642075-72-7
      642075-74-9 642075-75-0 642075-76-1
      642075-77-2
      RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
          (ribofuranosylcytidine methylvaline ester combined with other
antivirals for treatment of flaviviridae infections)
      640281-90-9 CAPLUS
     L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)
Absolute stereochemistry. Rotation (+).
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642075-50-1 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, 4-methylbenzenesulfonate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 104-15-4 CMF C7 HB 03 S

642075-51-2 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, methanesulfonate (salt) (9CI) (GA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 75-75-2

CMF C H4 03 S

642075-52-3 CAPLUS RN

L-Valine, 3'-ester with 2'-C-methylcytidine, acetate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9

CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 642075-53-4 CAPLUS CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxy-1,2,3-

McIntosh

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10/609,298
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propanetricarboxylate (salt) (9CI) (CA INDEX NAME)
      CRN 640281-90-9
      CMF C15 H24 N4 O6
Absolute stereochemistry. Rotation (+).
      CM
      CRN 77-92-9
CMF C6 H8 07
             CO2H
HO2C-CH2-C-CH2-CO2H
             ОН
     642075-54-5 CAPLUS
L-Valine, 3'-ester with 2'-C-methylcytidine, propanedicate (salt) (9CI)
(CA INDEX NAME)
      CM 1
      CRN 640281-90-9
CMF C15 H24 N4 O6
Absolute stereochemistry. Rotation (+).
      CM 2
      CRN 141-82-2
      CMF C3 H4 O4
HO2C-CH2-CO2H
    642075-55-6 CAPLUS
L-Valine, 3'-ester with 2'-C-methylcytidine, (2R,3R)-2,3-
dihydroxybutanedicate (salt) (9CI) (CA INDEX NAME)
```

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM

CRN 87-69-4 CMF C4 H6 O6

Absolute stereochemistry.

RN 642075-56-7 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, butanedioate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-15-6

CMF C4 H6 04

HO2C-CH2-CH2-CO2H

642075-57-8 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, benzoate (salt) (9CI) (CA INDEX NAME)

CM 1

McIntosh

10/609,298

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 65-85-0 CMF C7 H6 02

642075-58-9 CAPLUS

L-Ascorbic acid, compd. with L-valine 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 50-81-7

CMF C6 H8 O6

Absolute stereochemistry.

RN 642075-59-0 CAPLUS

McIntosh

10/609,298 CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-oxopentanedioate (salt) (9CI) (CA INDEX NAME) CM 1 CRN 640281-90-9 CMF C15 H24 N4 O6 Absolute stereochemistry. Rotation (+). H₂N CM CRN 328-50-7 CMF C5 H6 05 но2с-с-сн2-сн2-со2н 642075-60-3 CAPLUS RN 1-Valine, 3'-ester with 2'-C-methylcytidine, 2,3-dihydroxypropyl phosphate (salt) (9CI) (CA INDEX NAME) CM 1 CRN 640281-90-9 CMF C15 H24 N4 O6 Absolute stereochemistry. Rotation (+). CM 2

CM 2 CRN 57-03-4 CMF C3 H9 06 P

OH HO-CH2-CH-CH2-OPO3H2

RN 642075-61-4 CAPLUS CN L-Valine, 3'-ester with 2'-C-methylcytidine, formate (salt) (9CI) (CA INDEX NAME)

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10/609,298
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CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 64-18-6 CMF C H2 O2

о--- сн-- он

RN 642075-62-5 CAPLUS L-Valine, 3'-ester with 2'-C-methyloytidine, (2E)-2-butenedicate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-17-8 CMF C4 H4 O4

Double bond geometry as shown.

642075-63-6 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, propanoate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM

CRN 79-09-4 CMF C3 H6 O2

642075-64-7 CAPLUS RN

L-Valine, 3'-ester with 2'-C-methylcytidine, hydroxyacetate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9

CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 79-14-1 CMF C2 H4 03

но-с-сн2-он

642075-65-8 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxypropanoate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

McIntosh

CM

CRN 144-62-7 CMF C2 H2 O4

642075-68-1 CAPLUS L-Valine, 3'-deter with 2'-C-methylcytidine, (2Z)-2-butenedicate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

642075-69-2 CAPLUS

L-Valine, 3'-ester with 2'-C-methyloytidine, 2-hydroxybenzoate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

CM

CRN 69-72-7 CMF C7 H6 03

CO2H

642075-70-5 CAPLUS L-Valine, 3'-ester with 2'-C-methyloytidine, sulfate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 7664-93-9 CMF H2 O4 S

642075-71-6 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, nitrate (salt) (9CI) (CA INDEX NAME) CN

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

CM 2

CRN 7697-37-2 CMF H N 03

о— N— он

RN 642075-72-7 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, carbonate (salt) (9CI) (CA INDEX NAME)

CM 1 CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 463-79-6 CMF C H2 03

0 |-HO-C-OH

RN 642075-74-9 CAPLUS CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrobromide (9CI) (CA INDEX NAME)

10/609,298

●x HBr

642075-75-0 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydriodide (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●x HI

642075-76-1 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, carbonate (2:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 463-79-6 CMF C H2 03

642075-77-2 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, phosphate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9

CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 7664-38-2 CMF H3 04 P

640281-90-9P

National (Prig mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Blodgical study); PREP (Preparation); USES (Uses) (ribofuranosyloytidine methylvaline ester for treatment of flaviviridae infections) 640281-90-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

RN

CN

Absolute stereochemistry. Rotation (+).

642075-41-0

RL: RCT (Reactant); RACT (Reactant or reagent) (ribofuranosylcytidine methylvaline ester for treatment of flaviviridae infections)

RN

642075-41-0 CAPLUS Cytidihe, 2'-C-methyl-, 2',3',5'-tris(benzeneacetate) (9CI) (CA INDEX NAME)

20724-73-69 640725-70-89 642075-42-19 642075-43-29 642075-44-39 642075-48-79 RI: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (Reductant of respect)
(risofurancey)-dytidine methylvaline ester for treatment of
flaviviridae infections)
20724-73-6 CAPLUS
Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN

640725-70-8 CAPLUS L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methyleytidine (CA INDEX NAME)

Absolute stereochemistry.

RN

642075-42-1 CAPLUS Cytidine, N-[(dimethylamino)methylene]-2'-C-methyl- (CA INDEX NAME) CN

Absolute stereochemistry. Double bond geometry unknown.

RN

642075-43-2 CAPLUS Cytidine, N-[(dimethylamino)methylene]-5'-O-[(1,1-dimethylathyl)diphenylsilyl]-2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

642075-44-3 CAPLUS L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 5'-o-[(1,1-dimethylethyl)diphenylsilyl]-N-[(dimethylamino)methylene]-2'-C-methylocyldine (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

642075-48-7 CAPLUS Cytidine, 2'-0-methyl-N-(phenylacetyl)-, 2',3',5'-tris(benzeneacetate) (9CI) (CA INDEX NAME)

640725-71-9P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(ribofuranosylcytidine methylvaline ester for treatment of flaviviridae infections)

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HCl

- ANSWER 53 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:2898 CAPLUS DN 140:42424
 - Preparation of nucleoside derivatives as inhibitors of RNA-dependent RNA
- They are not the state of the s IN
- PA
- so
- DI Patent
- LA English

FAN.	FAN.CNT 1																	
	PATENT	KIND DATE			APPLICATION NO.						DATE							
PI	WO 2004	VO 2004000858			A2 20031231			WO 2003-US19172						20030617				
	WO 2004	2004000858			A3 20050512													
	W:	AE, A	AG, AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,		
		CO, 0	CR, CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
		GM, E	HR, HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KR,	KZ,	LC,	LK,	LR,	LS,		
		LT, I	LU, LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,	PG,		
		PH, PL, PT,		RO,	RU,	sc,	SD,	SE,	SG,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,		
		TZ, t	JA, UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW							
	RW:	GH, C	GM, KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,		
		KG, E	KZ, MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,		
		FI, E	TR, GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,		
		BF, E	BJ, CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG		
	CA 2488	CA 2488534			A1 20031231				CA 2003-2488534						20030617			
	AU 2003	AU 2003269890				2004	0106	AU 2003-269890						20030617				

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1851421 A2 20050713 EP 2003-751777 20030617
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, II, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, GK
       EP 1551421
                                                                  JP 2004-515870
US 2006-517295
       JP 2005530843
                                       T
                                                                                                      20030617
       US 20070004669
                                       A1
                                                20070104
                                                                                                      20060615
PRAI US 2002-390579P
                                       P
                                                20020621
       WO 2003-US19172
                                       ta
                                                20030617
      MARPAT 140:42424
```



AB The present invention provides nucleoside compds. I, wherein B is nucleobase; R1 is fluoromethyl, difluoromethyl, trifluoromethyl; R2 is H, F, amino, OH, SH, alkoxy, alkylcarbonyloxy, alkyl; R3 and R4 are independently H, Cn, N3, halogen, OH, SH, amino, alkoxy, alkylcarconyloxy, alkenyl, alkynyl; R5 is H, alkylcarbonyl, P309H4, P206H3, phosphophonyl; R6 and R7 independently H, Me, hydroxymethyl, fluoromethyl; and certain derivs. thereof which are inhibitors of RNA-dependent RNA viral polymerase. These compds. are inhibitors of RNA-dependent RNA viral replication and are useful for the treatment of RNA-dependent RNA viral infection. They are particularly useful as inhibitors of hepatitis C virus (HCV) NS5B polymerase, as inhibitors of HCV replication, and/or for the treatment of hepatitis C infection. The invention also describes pharmaceutical compns. containing such nucleoside compds. alone or in combination with other agents active against RNA-dependent RNA viral infection, in particular BCV infection. Also disclosed are methods of inhibiting RNA-dependent RNA polymerase, inhibiting RNA-dependent RNA viral replication, and/or treating RNA-dependent RNA viral infection with the nucleoside compds. of the present invention. Thus, 2-amino-9-(2-C-fluoromethyl- β -D-ribofuranosyl)-3,9-dihydropurin-6-one was prepared and tested as inhibitor of RNA-dependent RNA viral polymerase. Title compds. tested in the HCV NS5B polymerase assay exhibited IC50's less than 100 umol. 510765-51-2P 636581-91-4P 636581-92-5P 636581-93-6P 636582-01-9P 636582-02-0P 636582-03-1P 636582-04-2P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(Uses)
(preparation of nucleoside derivs. as inhibitors of RNA-dependent RNA viral polymerase)

RN 510765-51-2 CAPLUS CN Uridine, 2'-C-(fluoromethyl)- (CA INDEX NAME)

CN Offdire, 2 -C-(fidofomethy)- (CA IN

Absolute stereochemistry.

RN 636581-91-4 CAPLUS CN Cytidine, 2'-C-(fluoromethyl)- (9CI) (CA INDEX NAME) Absolute stereochemistry.

RN

636581-92-5 CAPLUS Cytidine, 2'-C-(fluoromethyl)-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

636581-93-6 CAPLUS Uridine, 2'-C-(fluoromethyl)-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

636582-01-9 CAPLUS

Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-(fluoromethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

636582-02-0 CAPLUS Cytidine 5 -(tetrahydrogen triphosphate), 2 -C-(fluoromethyl)-5-methyl-(SCI) (CA INDEX MAME)

636582-03-1 CAPLUS Uridine 5'-(tetrahydrogen triphosphate), 2'-C-(fluoromethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

636582-04-2 CAPLUS Uridine 5'-(tetrahydrogen triphosphate), 2'-C-(fluoromethyl)-5-methyl-(9CI) (CA INDEX NAME) CN

- ANSWER 54 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2003:892793 CAPLUS
- DN 139:365176
- Preparation of nucleoside derivatives for treating hepatitis
- C virus infection
- IN Roberts, Christopher Don; Dyatkina, Natalia B.; Keicher, Jesse D.; Liehr, Roberts, office the second of the second of
- PA
- so
- DT Patent
- English FAN.CNT 1

	PATENT	NO.		KIND DATE			APPLICATION NO.							DATE		
PI	WO 200	309329	A2	A2 20031113			WO 2003-US14237						20030506			
	WO 200	309329	A3	A3 20040318												
	W:	AE,	AG, AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
		CO,	CR, CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR, HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
		LS,	LT, LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
		PL,	PT, RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,
		UA,	UG, US,	UZ.	VC.	VN.	YU.	ZA.	ZM.	ZW						

```
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
              FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF,
              BJ, CF, CG, CI,
                              CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     CB 2484921
                           Al
                                               CA 2003-2484921
                                                                        20030506
     AU 2003232071
                           A1
                                               AU 2003-232071
                                                                        20030506
     US 20040063658
                                  20040401
                                               US 2003-431631
                                                                       20030506
                           A1
                                              EP 2003-747674
     EP 1501850
                           A2
                                                                       20030506
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
109581 A 20050329 BR 2003-9581
     BR 2003009581
                                               CN 2003-810239
     CN 1653077
                                                                        20030506
                                               JP 2004-501429
     JP 2005530759
                                                                        20030506
     NZ 536123
                                               NZ 2003-536123
                                                                        20030506
     MX 2004PA10983
                                  20050214
                                              MX 2004-PA10983
                                                                        20041105
                           ъ
                                              NO 2004-5247
     NO 2004005247
                           A
                                  20041130
                                                                       20041130
PRAI US 2002-378624P
                           p
     US 2002-392871P
                                  20020628
     WO 2003-US14237
                                  20030506
     MARPAT 139:365176
GΙ
```

AB Nucleosides I-III, wherein R and RI are independently H, alkyl, alkenyl, alkynyl, provided that R and RI are not both H; R2 is alkyl, cycloally, alkenyl, provided that R and RI are not both H; R2 is alkyl, cycloally, alkenyl, alkonyl, acylamino, guanddino, amidino, thioacylamino, OH, alkonyl, halo, nitro, aryl, heteroaryl, substituted amine; W is H, phosphate, phosphonate, acyl, alkyl, sulfonate, lipid, amino acid, sugar residue, peptide, cholesterol; W is H, halo, alkyl, univoitived amine; Y is H, halo, CH, alkylthio, substituted amine; Z is H, halo, CH, alkyl, substituted amine; I is nucleobase, were prepared as HCV RNA polymerase inhibitors and for treating hepatitis C virus infections. Thus, 2-(4-amino-pyrrolo), 2-c)cyridin-1-yl)-5-hydroxymethyl-1-methyltetrahydro-furan-3, 4-diol was prepared for treating for formulation such sublet, capital, Different kind suppository formulation are reported.

3148-85-41 high-10-64-3p 62380-51-2P

622380-52-3P 622380-56-7P 622380-57-8P 622380-59-0P 622380-60-3P 622380-61-4P

622380-59-0P 622380-60-3P 622380-61-4P 622380-89-6P 622380-90-9P 622381-09-3P

622381-10-6P RL: TM (Industrial manufacture); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of nucleoside derivs, for treating hepatitis

C virus infection)
31448-54-1 CAPLUS
Uridine, 2'-C-methyl- (CA INDEX NAME) CN

Absolute stereochemistry. Rotation (+).

119410-84-3 CAPLUS CN Uridine, 5-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

622380-51-2 CAPLUS

Cytidine, N-(2-amino-2-oxoethyl)-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

622380-52-3 CAPLUS
Pyriddinium, 1-(1/1,2-dihydro-1-(2-C-methyl-β-D-ribofuranosyl)-2-oxo-4pyrimidinyl aminojnethylj- (CA INDEX NAME)

Absolute stereochemistry.

McIntosh

10/609,298

RN

622380-56-7 CAPLUS (Sytidine, N-(2-amino-1-(3H-indol-3-ylmethyl)-2-oxoethyl]-2'-C-methyl-(9C1) (GA INDEX NAME)

Absolute stereochemistry.

622380-57-8 CAPLUS Uridine, 2'-C-methyl-, 4-[(pentafluorophenyl)hydrazone] (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- RN 622380-59-0 CAPLUS CN Cytidine, N-[2-(3H-indol-3-yl)ethyl]-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

622380-60-3 CAPLUS

Cytidine, N-(2-aminoethyl)-2'-C-methyl- (9CI) (CA INDEX NAME)

RN 622380-61-4 CAPLUS CN Cytidine, N-[1-(aminocarbonyl)-2-methylpropyl]-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 622380-89-6 CAPLUS

CN Furo[2,3-d]pyrimidin-2(1H)-one, 3,7a-dihydro-6-methyl-3-(2-C-methyl-β-D-ribofuranosyl)- (CA INDEX NAME)

Absolute stereochemistry.

RN 622380-90-9 CAPLUS

Furo[2,3-d]pyrimidin-2(1H)-one, 3,5,6,7a-tetrahydro-3-(2-C-methyl-β-D-ribofuranosyl)- (CA INDEX NAME)

Absolute stereochemistry.

RN 622381-09-3 CAPLUS

CN Cytidine, N-cyclopropy1-2'-C-methyl- (9CI) (CA INDEX NAME)

RN 622381-10-6 CAPLUS

Uridine, 2'-C-methyl-, 4-hydrazone (9CI) (CA INDEX NAME)

ANSWER 55 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN

Absolute stereochemistry.

2003:591195 CAPLUS

139:133789

```
Preparation of sugar modified nucleosides as antiviral agents
Hong, Zhi; An, Haoyun; Ding, Yili; Girardet, Jean-luc; Zhong, Weidong
IN
      Ribapharm Inc., USA
PCT Int. Appl., 33 pp.
CODEN: PIXXD2
PA
so
DT
       Patent
1.1
       English
FAN.CNT 4
                                                             APPLICATION NO.
       PATENT NO.
                                  KIND
                                            DATE
                                                                                               DATE
                                   ----
PT
      WO 2003062255
                                    A2
                                                             WO 2002-US231556
                                                                                               20021002
      WO 2003062255
                                    A3
                                             20060908
            W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                  CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, UF, KE, KG, KF, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MM, MX, MD, NO, NZ, OM, FH,
                  PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
                  UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
            RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, II, LU, MC, NL, FT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

20050914 EP 2002-776103

20021002

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, US 20070032448 A1 200700208 US 2008-535742 20060925 FRAI US 2002-350296P P 20020117 US 2002-391800P P 200201626

A2

WO 2002-US31556 W 2

EP 1572705

GI

1.6

AN

DN

- Various 2'-modified nucleoside analogs I and II wherein X is NH2, NHMe, NMe2, OMe, SMe, and corresponding prodrugs are provided, and particularly contemplated methods of use include use as antiviral agents, and especially as antiviral agents against HCV.
- antivara agents over the second of the secon
- (Biological study); USES (Uses)
 (preparation of sugar modified nucleosides as antiviral agents)
- 20724-73-6 CAPLUS Cytidine, 2'-C-methyl- (CA INDEX NAME) RN

Absolute stereochemistry.

RN 31448-54-1 CAPLUS Uridine, 2'-C-methyl- (CA INDEX NAME) CN

Absolute stereochemistry. Rotation (+).

RN 119410-84-3 CAPLUS

Uridine, 5-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

10/609,298

Absolute stereochemistry.

RN 565451-07-2 CAPLUS CN Cytidine, 5-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

RN 565451-08-3 CAPLUS CN Cytidine, N-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 565451-09-4 CAPLUS CN Cytidine, N,N-dimethyl-2'-C-methyl- (9CI) (CA INDEX NAME) Absolute stereochemistry.

RN 565451-10-7 CAPLUS CN Cytidine, N,5-dimethyl-2'-C-methyl- (9CI) (CA INDEX NAME) Absolute stereochemistry.

565451-11-8 CAPLUS CN Cytidine, N,N,5-trimethyl-2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- ANSWER 56 OF 58 CAPLUS COPYRIGHT 2008 ACS on SIN
- AN 2002:555629 CAPLUS
- 137:125359 DN
- Preparation of nucleoside derivatives as inhibitors of RNA-dependent RNA TT viral polymerase
- Carroll, Steven S.; Lafemina, Robert L.; Hall, Dawn L.; Himmelberger, Amy L.; Kuo, Lawrence C.; Maccoss, Malcolm; Olsen, David B.; Rutkowski, Carrie A.; Tomassini, Joanne E.; An, Haoyun; Bhat, Balkrishen; Bhat, Neelima; Cook, Phillip Dan; Eldrup, Anne B.; Guinosso, Charles J.; Prhavc, Marija; Prakash, Thazha P. PA
 - Merck & Co., Inc., USA; Isis Pharmaceuticals, Inc. PCT Int. Appl., 235 pp.
- CODEN: PIXXD2
- Patent
- English LA
- FAN.CNT 2 PATENT NO. KIND DATE APPLICATION NO. DATE ΡI WO 2002057425 A2 WO 2002-US1531 20020118 WO 2002057425 A3 20050421 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MM, MM, MX, NO, NZ, OK, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG CA 2433878 Al CA 2002-2433878 AU 2002-243600 AU 2002243600 A1 20020118 AU 2002243600 B2 US 20020147160 US 2002-52318 20020118 A1 US 6777395 B2 20040817 CN 1498221 20040519 CN 2002-806977 20020118 JP 2004532184 JP 2002-558479 EP 1539188 A2 20050615 EP 2002-709095 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR 371 A1 20061004 EP 2006-76021

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

20020118

EP 1707571

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IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
    ES 2278009
                                             ES 2002-709299
                                 20070801
     TW 261056
                                 20060901
                                              TW 2002-91100893
                                                                      20020121
    US 20040072788
                          2.1
                                             US 2003-431657
                                                                      20030507
     ZA 2003005078
                          А
                                             ZA 2003-5078
                                                                      20030630
     US 20040067901
                          A1
                                             US 2003-688691
                                                                      20031017
    US 7125855
                          B2
                                 20061024
     tts 20040110717
                                             HS 2004-250873
                                                                      20040116
                          2.1
     US 7105499
                          B2
                                 20060912
     US 20050272676
                                             US 2005-200499
                          A1
     US 20060205686
                          A1
                                 20060914
                                             US 2005-236224
                                                                      20050927
                           21
                                 20061123
                                             US 2006-496338
                                                                      20060731
     US 7202224
                          32
                                 20070410
     US 20070275912
                                 20071129
                                             US 2006-643464
                          A1
     JP 2007224045
                                 20070906
                                             JP 2007-115345
                          a
PRAI US 2001-263313F
     US 2001-282069P
                          P
                                 20010406
     US 2001-299320P
                                 20010619
     US 2001-344528P
                          P
     EP 2002-709299
                          A3
     JP 2002-558479
                          A3
     US 2002-52318
                           АЗ
     WO 2002-US1531
                          w
     US 2003-431657
                          В1
     US 2003-688691
                          A1
     US 2005-200499
                          B1
                                 20050809
    MARPAT 137:125359
```

The present invention provides the preparation of nucleoside compds. I, wherein AR B is nucleobase, Y is H, alkylcarbonyl, phosphate; R1 is H, alkenyl, alkynyl, alkyl; R2 and R3 are independently H, OH, halogen, alkyl, alkoxy, alkenyloxy, alkylthio, alkylcarbonyloxy, aryloxycrbonyl, azido, amino, alkylamino; R1 and R2 together with the carbon atom to which they are attached form a 3- to 6-membered heterocycle; R4 is H, OH, SH, NH2, alkylamino, cycloalkylamino, halogen, alkyl, alkoxy, CF3; R5 and R6 are independently H, hydroxymethyl, Me, fluoromethyl; and certain derivs.

thereof which are inhibitors of RNA-dependent RNA viral polymerase. These compds. are inhibitors of RNA-dependent RNA viral replication and are useful for the treatment of RNA-dependent RNA viral infection. They are particularly useful as inhibitors of hepatitis C virus (HCV) NS5B polymerase, as inhibitors of HCV replication, and/or for the treatment of hepatitis C infection. The invention also describes pharmaceutical compns. containing such nucleoside compds. alone or in combination with other agents active against RNA-dependent RNA viral infection, in particular HCV infection. Also disclosed are methods of inhibiting RNA-dependent RNA polymerase, inhibiting RNA-dependent RNA viral replication, and/or treating RNA-dependent RNA viral infection with the nucleoside compds. of the present invention. Thus, 4-amino-1-(2-C-methyl-β-Dribofuranosyl)-1H-pyrazolo[3,4-d]pyrimidine was prepared as inhibitors of RNA-dependent RNA viral polymerase. Representative compds. tested in the HCV NS5B polymerase assay exhibited IC's less than 100 μM . The compds. of the present invention were also evaluated for their ability to affect the replication of Hepatitis C Virus RNA in cultured hepatoma (HuH-7) cells containing a sub-genomic HCV Replicon. 20724-73-6P 114262-49-6P 374750-28-4P 444019-82-3P 444020-83-1P 444022-03-1P

RL: IMF (Industrial manufacture); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study);

(preparation of nucleoside derivs, as inhibitors of RNA-dependent human RNA

McIntosh

PREP (Preparation); USES (Uses)

viral polymerase) 20724-73-6 CAPLUS Cytidine, 2'-C-methyl- (CA INDEX NAME) CN

Absolute stereochemistry.

114262-49-6 CAPLUS

2,4(1H,3H)-Pyrimidinedione, 1-(2-C-methyl-β-D-arabinofuranosyl)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 374750-28-4 CAPLUS

Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 444019-82-3 CAPLUS

2,4(18,38)-Pyrimidinedione, 1-[5-0-[hydroxy[[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-2-C-methyl- β -D-arabinofuranosyl]- (CA INDEX NAME)

Absolute stereochemistry.

444020-83-1 CAPLUS 5'-Cytidylic acid, 2'-C-methyl-, bis[[[(1-methylethoxy)carbonyl]oxy]methyl | ester (SCI) (CA INDEX NAME)

444022-03-1 CAPLUS

2,4(1H,3H)-Byrimidinedione, 1-[2-C-methyl-5-0-[7-methyl-1-[[[(1-methylethoxy]carbonyl]oxy]methoxy]-1-oxido-5-oxo-2,4,6-trioxa-1-phosphaoct-1-yl]-[8-D-arabinofuranosyl]-[901] (CA INDEX NAME)

- ANSWER 57 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN 2001:886155 CAPLUS 1.6
- AN
- DN 136:590 TI
- Methods and compositions using modified nucleosides for treating
- TN
- Flaviviruses and pestiviruses
 Sommadossi, Jean-Pierre; Lacolla, Paolo
 Novirio Pharmaceuticals Limited, Cayman I.; Universita Degli Studi Di PA
- Cagliari PCT Int. Appl., 302 pp. CODEN: PIXXD2 so
- DT Patent

	Enq.	glish 2																	
	PA:	KIND DATE				APPL	ICAT		D.	ATE									
PI	WO	2001092282			A2 20011206				WO 2	001-		20010523							
	WO	2001092282			A3 20020502														
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			LS,	LT.	LU,	LV,	MA,	MD,	MG.	MK,	MN,	MW.	MX,	MZ.	NO.	NZ,	PL,	PT,	
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	UZ. VN. YU.																		
		RW:	GH,	GM.	KE.	LS.	MW.	MZ.	SD.	SL.	SZ.	TZ.	UG,	ZW.	AT.	BE,	CH.	CY,	
			DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	
			BJ.	CF.	CG.	CI.	CM.	GA.	GN.	GW.	ML.	MR.	NE.	SN.	TD.	TG			
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	EP	1294	735			A2 20030326				EP 2	001-	9521	20010523						
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
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	US	2003	0060	400		A1		2003	0327		US 2	001-	8638	16		20010523			
	US	6812	219			B2		2004	1102										
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	JP	2004	5106	98		T		2004	0408		JP 2	002-	5008	95		20010523			
	NZ	Z 536570			A		2006	0831		NZ 2	001-	5365	70		2	0010	523		
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		1727	A	20070430		2002-2705	20010523
	CN	101099745	A	20080109	CN	2007-10089609	20010523
	NZ	547204	A	20080131	NZ	2001-547204	20010523
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	US	7148206	B2	20061212			
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		7163929	B2	20070116			
		20070037773	A1	20070215		2006-527124	20060925
		2007202602	A1	20070719		2007-202602	20070607
	KR	2008021797	A	20080307	KR	2008-701618	20080121
PRAI		2000-207674P	P	20000526			
	US	2001-283276P	P	20010411			
		2001-813182	A3	20010523			
	EP	2001-952131	A3	20010523			
	US	2001-863816	A3	20010523			
		2001-US16687	W	20010523			
	KR	2002-715794	A3	20021122			
		2003-602135	A1	20030620			
os	MAI	RPAT 136:590					

A method and composition are provided for treating a host infected with

flavivirus or pestivirus, comprising administering an effective amount of a 1', 2' or 3'-modified nucleoside or a pharmaceutically acceptable salt or prodrug thereof. 20724-73-6 31448-54-1 119410-84-3

RI: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nucleoside derivs. for treating flaviviruses and

pestiviruses)
20724-73-6 CAPLUS
Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

31448-54-1 CAPLUS Uridine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

119410-84-3 CAPLUS

Uridine, 5-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

125911-76-4 374750-28-4

RL: BSU (Biological study, unclassified); PKT (Pharmacokinetics); BIOL (Biological study)

(nucleoside derivs. for treating flaviviruses and pestiviruses

125911-76-4 CAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

374750-28-4 CAPLUS

Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (CA INDEX NAME) CN Absolute stereochemistry.

- ANSWER 58 OF 58 CAPLUS COPYRIGHT 2008 ACS on STN
- 2001:868467 CAPLUS
- 136:6296 DN
- Preparation of antiviral nucleosides and methods for treating hepatitis C virus
- Sommadossi, Jean-Pierre; Lacolla, Paulo Novirio Pharmaceuticals Limited, Cayman I.; Universita degli Studi di
- PA
- Cagliari PCT Int. Appl., 296 pp. so
- CODEN: PIXXD2
- Patent DT
- LA
- English FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001090121	A2	20011129	WO 2001-US16671	20010523
	MO 2001090121	2.2	20020502		

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RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
              UZ, VN, YU, ZA, ZW
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                            D.
                                               EP 2006-75216
     EP 1669364
                            A2
                                   20060614
     EP 1669364
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PRAI US 2000-206585F
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                            АЗ
                                   20010523
     EP 2001-941564
                            A3
     NZ 2001-522863
                            73
     US 2001-864078
                            A1
     WO 2001-US16671
                            w
     KR 2002-715790
                            А3
     NO 2002-5627
                            Α
      IN 2002-DN1184
                            A3
os
     MARPAT 136:6296
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AB A method and composition for treating a host infected with hepatitis C comprising administering an effective hepatitis

C treatment amount of a described 1'-, 2'- or 3'-modified

nucleosides I, wherein : R1-R3 and R are independently H, phosphate (including mono, di- or triphosphate and a stabilized phosphate prodrug); acyl; alkyl; sulfonate ester including alkyl or arylalkyl sulfonyl including methanesulfonyl and benzyl, wherein the Ph group is optionally substituted with one or more substituents as described in the definition of aryl given herein; a lipid, including a phospholipid; an amino acid; a carbohydrate; a peptide; a cholesterol; or other pharmaceutically acceptable leaving group which when administered in vivo is capable of providing a compound wherein R1-R3 are independently H or phosphate; Y is hydrogen, bromo, chloro, fluoro, iodo, OR4, NR4R5 or SR4; X1 and X2 are independently selected from the group consisting of H, straight chained, branched or cyclic alkyl, CO-alkyl, CO-arkyl, CO-alkoxyalkyl, chloro, bromo, fluoro, iodo, OR4, NR4R5 or SR4; and R4 and R5 are independently hydrogen, acyl, alkyl.or a pharmaceutically acceptable salt or prodrug thereof, is provided. Thus, I (R1-R3=X1=X2=H, Y=NH2) was prepared and tested in Cynomolgus monkeys as antiviral agent. Oral bioavailability in monkeys, bone human bone marrow toxicity (IC50 > 10 μ M), and mitochondrial toxicity, were reported . 20724-73-6P 31448-54-1P 119410-84-3P

125911-76-4P 374750-28-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); IMF (Industrial manufacture); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of antiviral nucleosides and methods for treating hepatitis C virus)

20724-73-6 CAPLUS RN Cytidine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

31448-54-1 CAPLUS RN Uridine, 2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

119410-84-3 CAPLUS

Uridine, 5-methyl-2'-C-methyl- (9CI) (CA INDEX NAME)

RN 125911-76-4 CAPLUS
CN Uridine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Absolute stereochemistry.

RN 374750-28-4 CAPLUS CN Cytidine 5'-(tetrahydrogen triphosphate), 2'-C-methyl- (CA INDEX NAME)

=> file reg COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 132.83 311.40 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -15.20 -15.20

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DICTIONARY FILE UPDATES: 6 JUN 2008 HIGHEST RN 1026208-38-7

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http://www.cas.org/support/stngen/stndoc/properties.html

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* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

4 ANSWERS

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Structure attributes must be viewed using STN Express query preparation.

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SAMPLE SEARCH INITIATED 18:14:14 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -2232 TO ITERATE

89.6% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE** BATCH **COMPLETE** PROJECTED ITERATIONS: 41806 TO 47474 PROJECTED ANSWERS: 4 TO 215

4 SEA SSS SAM L7

FULL SEARCH INITIATED 18:14:19 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 44523 TO ITERATE

100.0% PROCESSED 44523 ITERATIONS 171 ANSWERS SEARCH TIME: 00.00.01

171 SEA SSS FUL L7

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CA SUBSCRIBER PRICE

=> s 17 full

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McIntosh

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http://www.cas.org/legal/infopolicy.html
=> s 19
L10
            74 1.9
=> s 110 and (flavivirus or pestivirus or flaviviridae or how or hepatitis c)
          1747 FLAVIVIRUS
           864 FLAVIVIRUSES
          2025 FLAVIVIRUS
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           501 PESTIVIRUS
           266 PESTIVIRUSES
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L11
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L11 ANSWER 1 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
     2008:562068 CAPLUS
AN
DN
     148:509492
     The hepatitis C virus replicon presents a higher
     barrier to resistance to nucleoside analogs than to nonnucleoside
     polymerase or protease inhibitors
AU
     McCown, Matthew F.; Rajyaguru, Sonal; Le Pogam, Sophie; Ali, Samir; Jiang,
     Wen-Rong; Kang, Hyunsoon; Symons, Julian; Cammack, Nick; Najera, Isabel
    Department of HCV Biology, Virology Disease Biology Area, Roche Palo Alto LLC, Palo Alto, CA, 94304, USA
so
    Antimicrobial Agents and Chemotherapy (2008), 52(5), 1604-1612
     CODEN: AMACCQ; ISSN: 0066-4804
PR
    American Society for Microbiology
LA
    English
AR
     Specific inhibitors of hepatitis C virus (HCV
     ) replication that target the NS3/4A protease (e.g., VX-950) or the NS5B polymerase (e.g., R1479/R1626, PSI-6130/R7128, NM107/NM283, and
     HCV-796) have advanced into clin. development. Treatment of
     patients with VX-950 or HCV-796 rapidly selected for
     drug-resistant variants after a 14-day monotherapy treatment period.
     However, no viral resistance was identified after monotherapy with R1626
     (prodrug of R1479) or NM283 (prodrug of NM107) after 14 days of
     monotherapy. Based upon the rapid selection of resistance to the protease
     and nonnucleoside inhibitors during clin, trials and the lack of selection
     of resistance to the nucleoside inhibitors, we used the replicon system to
     determine whether nucleoside inhibitors demonstrate a higher genetic barrier to
     resistance than protesse and nonnucleoside inhibitors. Treatment of
     replicon cells with nucleoside inhibitors at 10 and 15 times the 50%
     effective concentration resulted in clearance of the replicon, while treatment
     with a nonnucleoside or protease inhibitor selected resistant colonies.
     In combination, the presence of a nucleoside inhibitor reduced the
     frequency of colonies resistant to the other classes of inhibitors.
     results indicate that the HCV replicon presents a higher barrier
     to the selection of resistance to nucleoside inhibitors than to
    nonnucleoside or protease inhibitors. Furthermore, the combination of a
     nonnucleoside or protesse inhibitor with a nucleoside polymerase inhibitor
```

could have a clear clin, benefit through the delay of resistance emergence.

640725-71-9, NM283

RE: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(hepatitis C virus replicon presents a higher

barrier to resistance to nucleoside analogs than to nonnucleoside polymerase or protease inhibitors) 640725-71-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HC1

RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN CAPLUS AN

2008:416659 DM

TT

Compositions of immunostimulatory oligonucleotides as Toll-like receptor ligands and antiviral agents for therapeutic administration Vollmer, Jorg; Jurk, Marion; Uhlmann, Eugen; Debelak, Harald; Bratzler, Robert L; Vicari, Alain

PA Coley Pharmaceutical Group, Inc., USA; Coley Pharmaceutical G.m.b.H.;

Coley Pharmaceutical Group, Ltd.

PCT Int. Appl., 89pp. CODEN: PIXXD2

DТ Patent

LA	Eng	lish																
FAN.	FAN.CNT 1																	
	PATENT NO.						KIND DATE			APPLICATION NO.						DATE		
PI	WO 2008039538					A2 20080403			WO 2007-US21030						20070927			
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		RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,
			IS,	IT,	LT,	LU,	LV,	MC,	MT,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,
			BJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG,	BW,
			GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,
			BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM									
PRA:	US.	2006	-847	408P		P		2006	0927									

The invention relates to methods and products for the treatment of viral infection using a combination of antiviral agents and Toll-like receptor (TLR) ligands. The TLR ligands comprise immunostimulatory oligonucleotides, preferably containing modifications selected from 8-oxo-rG, 8-bromo-dG, 8-bromo-dA, and isatoribine (Immunosine) with a 5'-5' linkage. The 8-modified guanine residues enhance immunostimulatory activity, particularly when present at the 5' end of the oligonucleotide. Combination of Ribavirin with an immunostimulatory CpG-containing oligonucleotide results in a decrease of interleukin-10 relative to interferon- α inducing activity. Further, Ribavirin and CpG

oligonucleotide improve survival in a mouse cancer model. 640281-90-9, Valopicitabine RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(compns. of immunostimulatory oligonucleotides as Toll-like receptor ligands and antiviral agents for therapeutic administration) 640281-90-9 CAPLUS RN

L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME) CN

Absolute stereochemistry. Rotation (+).

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L11 ANSWER 3 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
ΔN
      2008:352859 CAPLUS
DN
      148:394354
      Compositions and methods for treatment of viral diseases
      Johansen, Lisa M.; Owens, Christopher M.; Mawhinney, Christina; Chappell, Todd W.; Brown, Alexander T.; Frank, Michael G.; Altmeyer, Ralf
IN
```

PA Combinatorx (Singapore) Pre. Ltd., Singapore so PCT Int. Appl., 237pp.

CODEN: PIXXD2 Patent

LA English

FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE ----PI WO 2008033466 A2 20080320 WO 2007-US19932 20070913 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, II, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

PRAI US 2006-844463P P 20060914 US 2006-874061P 20061211

Based on the results of the authors screen identifying compds, and combinations of compds. having antiviral activity, the present invention features compns., methods, and kits useful in the treatment of viral In certain embodiments, the viral disease is caused by a single stranded RNA virus, a flaviviridae virus, or a hepatic virus. In particular embodiments, the viral disease is viral hepatitis (e.g., hepatitis A, hepatitis B, hepatitis C, hepatitis D, hepatitis E). Also featured are screening methods for identification of

novel compds, that may be used to treat a viral disease. 640281-90-9, Valopicitabine 640725-71-9, NM-283

1015079-99-8 1015080-00-8 1015080-23-5

1015080-56-4 1015080-58-6 1015080-59-7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(compns. and methods for treatment of viral diseases) 640281-90-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

640725-71-9 CAPLUS L-Valine, 3'-ester with 2'-C-methyloytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HC1

1015079-99-8 CAPLUS L-Valine, 31-ester with 2'-C-methylcytidine, hydrochloride (1:2), mixt. with (3R,5a5,6R,6as,9R,12S,12RR)-octahydro-3,6,9-trimethyl-3,12-epoxy-12H-pyrano(4,5-d)-1,2-benzodloxepin-10(5H)-one (CA INDEX NAME) CN

CM

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

●2 HCl

CM

CRN 63968-64-9 CMF C15 H22 O5

10/609.298

N 1015080-00-8 CAPLUS
N L-Valine, 3'-ester with 2'-C-methylogidine, hydrochloride (1:2), mixt. with 1,8,9-trihydroxy-3-methoxy-6H-benzofuro[3,2-c][1]benzopyran-6-one (CA INDEX NAME)

CM 1

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

● 2 HCl

CM

CRN 524-12-9 CMF C16 H10 07

RN 1015080-23-5 CAPLUS

NN 1-Valine, 31-ester with 2'-C-methyloytidine, hydrochloride (1:2), mixt. with (62)-N-[(4-hydroxy-3-methoxyphenyl)methyl]-8-methyl-6-nonenamide (CA INDEX NAME)

CM 1

CRN 640725-71-9

CMF C15 H24 N4 O6 . 2 Cl H

10/609,298

● 2 HC1

CM

CRN 404-86-4 CMF C18 H27 N O3

Double bond geometry as shown.

1015080-28-0 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2), mixt. with 3-[(3,5-dibromo-4-hydroxyphenyl)methylene]-1,3-dihydro-5-iodo-2H-indol-2-one (CA INDEX NAME)

CM 1

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

●2 HC1

CM

CRN 220904-83-6 CMF C15 H8 Br2 I N 02

1015080-31-5 CAPLUS L-VAline, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2), mixt. with 2,3,6'-Tetrahydro-9,10-dimethoxy-3-methyl-2-[(2,4,6-trimethylphenyl)imino|-4H-pyrimido[6,1-a]isoquinolin-4-one (CA INDEX NAME)

CM

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 Cl H

Absolute stereochemistry. Rotation (+).

2 HC1

CM

CRN 79855-88-2 CMF C24 H27 N3 O3

1015080-38-2 CAPLUS

L-Valine, 3'-ester with 2'-C-methyloytidine, hydrochloride (1:2), mixt. with 2,3-dihydro-2-hydroxy-4H-1-benzopyran-4-one (CA INDEX NAME)

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

10/609,298

●2 HC1

CM

CRN 57669-32-6 CMF C9 H8 O3

RN

1015080-56-4 CAPLUS L-Valine, 3'-ester with 2'-0-methyloytidine, hydrochloride (1:2), mixt. with 1:(4-fluorophenyl)-4-[4-hydroxy-4-(3-(trifluoromethyl)phenyl)-1-piperidinyl]-1-butanone (CA INDEX NAME)

CM

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

●2 HC1

CM 2

CRN 749-13-3 CMF C22 H23 F4 N O2

1015080-58-6 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2), mixt. with 3-(2-phenyldiazenyl)-2,6-pyridinediamine (CA INDEX NAME)

CM

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

●2 HC1

CM 2

CRN 94-78-0 CMF C11 H11 N5

1015080-59-7 CAPLUS

L-Valine, 3'-ester with 2'-C-methyloytidine, hydrochloride (1:2), mixt. with α,α,α -trifluorothymidine (CA INDEX NAME)

CRN 640725-71-9 CMF C15 H24 N4 O6 . 2 C1 H

Absolute stereochemistry. Rotation (+).

●2 HC1

CM

CRN 70-00-8 CMF C10 H11 F3 N2 O5

Absolute stereochemistry.

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L11 ANSWER 4 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
AN
    2008:90893 CAPLUS
DN
    148:192198
    Preparation of peptidomimetics as modulators of pharmacokinetic properties
```

of therapeutics by inhibiting cytochrome P450 monooxygenase Desai, Manoj C.; Hong, Allen Yu; Liu, Hongtao; Xu, Lianhong; Vivian, Randall W.

PA

Gilead Sciences, Inc., USA PCT Int. Appl., 346pp. CODEN: PIXXD2 so

DT Patent LA English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE WO 2008010921 A2 PT 20080124 WO 2007-US15604 20070706 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, FG, FH, FL, FT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MG, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, MI, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM US 20080108617 US 2007-825605 20070706 A1 PRAI US 2006-819315P P US 2006-832371P 20060721

US 2007-903228P MARPAT 148:192198

The invention is related to the preparation of R8YZ1[CONR1(CR2R2)m]nL1NR3CH[L3A AB. (L4Ar)p]CHR4L2CH[L3A(L4Ar)p]NR5COZ2XR9 [I; L1 = C(R6)2, CO, SO2, NHCO and derivs, OCO; R4, R6 = independently H, heteroalkyl, (un)substituted alkyl; 12 = a covalent bond, C(R6)2, CO; each L3 = independently a covalent bond, (un)substituted alkylene; each L4 = L3, O, CR2O, NH; each A = H, (un)substituted alkyl, aryl, heterocyclyl with the proviso that when A = H, p = 0; Z1, Z2 = independently O, NH and derivs.; <math>Y, X =independently heterocyclyl, heterocyclylalkyl; each Ar = independently (un) substituted (hetero) aryl; R1, R3, R5 = independently H, (un) substituted aryl/alkyl; each R2 = independently H, (un) substituted arylhetero/hydroxy/amino/alkyl, alkylene-CO2H, alkylene-CO-alkyl, etc.; R8, R9 are each one or more H's or substituents selected from Cl. CN, (un) substituted alkyl, aryl, heterocyclyl, m=1-2, n=0-1; each p=1 independently 0-1), their pharmaceutically acceptable salts, solvates and esters, and compns. containing them which improve the pharmacekinetics of a co-administered drug which is metabolized by cytochrome P 450 on-naminister ding which is meta-red by cyclohodie = 300 monoxygense. Thus, a multi-step synthesis using 2-ioopropyl-4- [(methylamino|methyl]-1,3-thiazole, (23)-2-amino-4-[(tert-butoxycarbonyl)amino|butanoic acid Me ester, amine II and (BrCR2CH2)20 was given for III. III inhibited CYP450 3A4 (IC50 = 80-150 nM), CYP450 2C9 (IC50 = 1,000-10,000 nM) and protease (EC50 > 20,000 nM in an anti HIV-1 cell culture assay). I alone or in combination with one or more addnl. therapeutic agents which are metabolized by cytochrome P 450 monocxygenase are useful for treating a viral infection, e.g. HIV (no data).

are userul for treating a viral insection, e.g. hiv (ho data), 640281-9-9, Valopicitabine 640728-71-9, Mm-2933
Ri: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (compds. as modulators of pharmacokinetic properties of therapeutic

(compds. as modulators of pharmacokinetic properties of therapeuti agents)

640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HC1

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L11 ANSWER 5 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
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2008:40914 CAPLUS AN 148:168504

DN

Preparation of purine and thiadeazapurine phosphonate derivatives as

Preparation of purise and thiasesapurine phosphonate derivatives modulators of toll-like sceeptor independent of the state IN

so

DT Patent

English

FAN.	CNT 1																
	PATENT	NO.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D	ATE	
						-											
PI	WO 2008	00555	5		A1		2008	0110		WO 2	007-	US15	615		21	0070	706
	W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BH,	BR,	BW,	BY,	BZ,	CA,
		CH,	CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DO,	DZ,	EC,	EE,	EG,	ES,	FI,
		GB,	GD,	GE,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,
		KM,	KN,	KP,	KR,	KZ,	LA,	LC,	LK,	LR,	LS,	LT,	LU,	LY,	MA,	MD,	ME,
		MG,	MK,	MN,	MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,
		PT,	RO,	RS,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	sv,	SY,	TJ,	TM,	TN,
		TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	ZW				
	RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,
		IS,	II,	LT,	LU,	LV,	MC,	MT,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,
		BJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG,	BW,
		GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,
		BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM									
	US 2008				A1		2008	0110		US 2	007-	8253	77		21	0070	706
PRAI	US 2006	-8194	90P				2006										
	US 2006	-8328	51P		P		2006	0724									
os	MARPAT	148:1	.6850	14													

GI

10/609.298

AB The present application provides for a compound I [Z = 0H, NHZ; XI = (un) supplicitude alkylene, alknylene, alknylene, carbooyclylene, provides of the compound of the composition of

II 640281-90-9, Valopicitables 640725-71-9, NN-283
Ri: PRC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(compds. as modulators of Toll-like receptor 7 useful in combination

(compds. as modulators of Toll-like receptor 7 useful in combination therapy and prevention of TLR7 activation-related diseases)
RN 640281-90-9 CAPUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 640725-71-9 CAPLUS

10/609,298

CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HC1

L11 ANSWER 6 OF 37 CAPLUS COPYRIGHT 2008 ACS on SIN

AN 2007:1332915 CAPLUS

DN 148:11439 TI 2'-C-Methyl-Ribofuranosyl Cytidine Prodrugs, Pharmaceutical Compositions and Uses Thereof

IN Gallop, Mark A.

PA USA SO U.S. Pat. Appl. Publ., 59pp.

CODEN: USXXCO DT Patent

LA English

GT

FAN.CNT 1 PATENT NO.

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	US 20070270374	A1	20071122	US 2007-752214	20070522
PRAI	US 2006-808360P	P	20060522		
OS	MARPAT 148:11439				

AB The present disclosure provides the preparation of 2'-C-mathyt-ribofuranosyl cytidine produys I, wherein Ri an R2 are independently R, acyl, acyloxyalkylcarbonyi, oxycarbonyi, substituted aninosanbonyi, R3 is substituted inine, substituted anine compast thereof to treat viral diseases such as hepatitis C. Thus, p. (4-allyloxycarbonyi approach (10 data) and tearly-ly-2-C mathibufurance was prepared (10 data). The propared in the record antivirial agent is selected from an interferon, ribavirin, interleukin, an N33 protease inhibitor, cysteine protease inhibitor, thanolidine derivative, thiazolidine, benarallide, phenathenequinone, a helicase inhibitor of REZ-dependent phosphorochicate oligodeoxyribonucleotides, inhibitor of REZ-dependent (17), (27),

957687-55-7P 957687-58-0P 957687-62-6P 957687-64-8P 957687-83-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(2'-C-methyl-ribofuranosyl cytidine prodrugs, pharmaceutical compns.

and uses thereof)

and used chereol)
937687-30-6 CAPLUS
1-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with
5'-0-[(1,1-dimethylethyl)dimethylsilyl]-N-(ethoxycarbonyl)-2'-Cmethylcytddine (CA INDEX NAME)

Absolute stereochemistry.

957687-34-2 CAPLUS

L-Valine, 3'-ester with N-(ethoxycarbonyl)-2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

957687-53-5 CAPLUS Cytidine, 2'-C-methyl-, 3'-(pentyl carbonate) (CA INDEX NAME)

Absolute stereochemistry.

957687-55-7 CAPLUS RN

Cytidine, 2'-C-methyl-N-[(pentyloxy)carbonyl]-, 3'-(pentyl carbonate) (CA INDEX NAME)

RN

957687-58-0 CAPLUS Cytidine, 2'-C-methyl-, 3',5'-bis(pentyl carbonate) (CA INDEX NAME)

Absolute stereochemistry.

957687-62-6 CAPLUS

Cytidine, 2'-C-methyl-N-[(pentyloxy)carbonyl]-, 3',5'-bis(pentyl carbonate) (CA INDEX NAME)

Absolute stereochemistry.

- 957687-64-8 CAPLUS
- Cytidine, 3',5'-bis-O-[(1,1-dimethylethyl)dimethylsilyl]-2'-C-methyl- (CA INDEX NAME)

Absolute stereochemistry.

957687-83-1 CAPLUS

Cytidine, 2'-C-methyl-N-[(pentyloxy)carbonyl]-, 3'-acetate (CA INDEX NAME 1

Absolute stereochemistry.

L11 ANSWER 7 OF 37 CAPLUS COPYRIGHT 2008 ACS on SIN 2007:1164795 CAPLUS

AN

DN 147:534049

2'-C-methyl branched pyrimidine ribonucleoside analogues: potent inhibitors of RNA virus replication

Benzaria, Samira; Bardiot, Dorothee; Bouisset, Tony; Counor, Clement; Rabeson, Celine; Pierra, Claire; Storer, Richard; Loi, Anna Giulia; Cadeddu, Alessandra; Mura, Massimo; Musiu, Chiara; Liuzzi, Michel; Loddo, Roberta; Bergelson, Svetlana; Bichko, Vadim; Bridges, Edward; Cretton-Scott, Erika; Mao, John; Sommadossi, Jean-Pierre; Seifer, Maria;

Standring, David; Tausek, Michele; Gosselin, Gilles; La Colla, Paolo Laboratoire Cooperatif Idenix-CNRS-Universite Montrellier II, Montrellier,

so Antiviral Chemistry & Chemotherapy (2007), 18(4), 225-242

CODEN: ACCHEH; ISSN: 0956-3202 PB International Medical Press, Ltd.

Journal

LA English

CASREACT 147:534049

RNA viruses are the agents of numerous wide-spread and often severe diseases. Their unique RNA-dependent RNA polymerase (RDRP) is essential for replication and, thus, constitutes a valid target for the development of selective chemotherapeutic agents. In this regard, the authors have investigated sugar-modified ribonucleoside analogs as potential inhibitors of the RDRP. Title compds. retain 'natural' pyrimidine bases, but possess a β -Me substituent at the 2'-position of the D- or L-ribose moiety. Evaluation against a broad range of RNA viruses, either single-stranded pos. (ssRNA+), single-stranded neg. (ssRNA-) or double-stranded (dsRNA), revealed potent activities for D-2'-C-methyl-cytidine and -uridine against ssRNA+, and dsRNA viruses. None of the L-enantiomers were active. Moreover, the 5'-triphosphates of the active D-enantiomers were found to inhibit the bovine virus diarrhea virus polymerase. Thus, the 2'-Me branching of natural pyrimidine ribonucleosides transforms physiol. mols. into potent, broad-spectrum antiviral agents that merit further

development. 23643-36-9P 957535-48-7P 957535-51-2P 957535-53-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (pyrimidine ribonucleoside analogs as potent inhibitors of RNA virus

replication)

23643-36-9 CAPLUS

2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-benzoyl-2-C-methyl-β-D-ribofuranosyl)- (9CI) (CA INDEX NAME)

10/609,298

RN 957835-48-7 CAPLUS
CN 2,4(1H,3H)-Pyrimidinedione, 5-methyl-1-(2,3,5-tri-0-benzoyl-2-C-methylB-L-ribofuranosyl)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 957535-51-2 CAPLUS
CN 2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-benzoyl-2-C-methyl-β-L-ribofuranoyyl)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 957535-53-4 CAPLUS
CN Uridine, 5-methyl-2'-C-methyl-, 2',3',5'-tribenzoate (CA INDEX NAME)

THERE ARE 73 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 8 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2007:1121524 CAPLUS

DN 147:407046

Preparation of nucleosides as Hepatitis C virus NS5b TI

polymerase inhibitors and antiviral agents via regioselective O-acylation reaction IN

PA

Sarma, Keshab Roche Palo Alto LLC, USA U.S. Pat. Appl. Publ., 15pp. SO

CODEN: USXXCO

Patent

FAN		1	
	70.70	AD TO A CAD	v

		ENT :				KIN	0	DATE			APPL						ATE	
PI	US	2007	0232	562		A1 A1		2007			US 2	007-	7329			2	0070	404
	no	W:	AE,	AG,		AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BH,	BR,	BW,	BY,	BZ,	CA,
			GD,	GΕ,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KM,
			MN,	MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,
		DAT.	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	ZW						
		ZW.	IS,	IT,	LT,	LU,	LV,	MC,	MT,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,
			GH,	GM,	KE,	LS,	MW,	GA, MZ,	NA,									
								IJ,										

PRAI US 2006-789491P P 20060404 OS CASREACT 147:407046; MARPAT 147:407046 GI

Nucleosides I, wherein Rl is Compds. having the formula I wherein Rl is C2-5 (un)-branched alkyl, C2-5 (un)-branched alkenyl, C3-5 cycloalkyl, C2-5 lower halo-alkyl, were prepared as Repatitis C

virus NS5b polymerase inhibitors. Also disclosed are compus, and methods for inhibiting hepatitis replication, processes for making the compds, and synthetic intermediates used in the process. Thus, nucleoside I.RCI [RI = C(0)EC] was prepared in 60% yield by regionsclettive O-acylation of I (RI = R) with propionyl obloride. Title compds, were tested in vivo a Bepatitis C virus NS5b polymerase inhibitors and antiviral agents (a dose of between 1.0 and 6.0 g per day is administered

antiviral agents (a dose of between 1.0 and 6.0 g per day is administered to the patient). Determination of pharmacokinetic parameters of title nucleosides in rate, is reported.

T S1131-56-3P 951131-58-1P 951131-60-5P R1: PAC (Pharmacological activity); PKT (Pharmacokinetice); GPN (Synthetic preparation); ITU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USEC (Uses)

(preparation of nucleosides as Hepatitis C virus NS5b polymerase inhibitors and antiviral agents via regioselective O-acylation reaction)

3 951131-56-9 CAPLUS Cytldine, 2'-C-methyl-, 3',5'-dipropanoate, hydrochloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.

HCl

RN 951131-58-1 CAPLUS
CN Cytidine, 2'-C-methyl-, 3',5'-bis(2-methylpropanoate), hydrochloride (1:1)
(CA INDEX NAME)

Absolute stereochemistry.

● HCl

RN 951131-60-5 CAPLUS CN Cytidine, 2'-C-methyl-, 3',5'-dipentanoate, hydrochloride (1:1) (CA INDEX NAME)

640281-90-9

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (preparation of nucleosides as Hepatitis C virus NS5b polymerase inhibitors and antiviral agents via regioselective O-acylation reaction)

RN 640281-90-9 CAPLUS CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

L11 ANSWER 9 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

Absolute stereochemistry. Rotation (+).

2007:1112011 CAPLUS 147:514184

New therapies for hepatitis C

AN DN

Nea Libergues A.; Hoofingle, Jay H. Liver Diseases Dranch, National Institute of Diabetes and Digestive and Kidney Diseases, National Institutes of Health, Betheeda, MD, USA Hepatology (Hoboker, NJ, United States) (2007), 46(3), 615-617 CODEN: HPTLD9; ISSN: 0270-9139 SO John Wiley & Sons, Inc. PB Journal; General Review LA English A review. The research Forestier et al. (2007) entitled "Antiviral activity of telaprevir (VX-950) and peginterferon alfa-2a in patients with hepatitis C" is reviewed with commentary and refs. Forestier and her coinvestigators from Saarland University Hospital, the University of Amsterdam, and Vertex Pharmaceutics describe the preliminary clin. results of a small phase lb trial of telaprevir. The report provides information on HCV RNA and alanine aminotransferase levels in 8 patients who received telaprevir alone, 8 who received telaprevir with peginterferon, and 4 who served as controls and received peginterferon alone for 2 wk. Telaprevir was then stopped, but the patients were offered a continuation of treatment with a combination of peginterferon and ribavirin until 48 wk and thus were provided the standard of care for chronic hepatitis C, genotype 1. Telaprevir led to a rapid decline in HCV RNA levels within 1-4 days. The

combination of peginterferon with telaprevir resulted in a similar early decline in viral levels, but importantly, the combination therapy was associated with an addnl., continuing decline after the first 4 days of

640281-90-9, Valopicitabine RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

treatment.

RN

(Biological study); USES (Uses) (valopicitabine, BILN-2061 showed greater toxicity hence were abandoned from usage by patient with hepatitis C)
640281-90-9 CAPLUS
L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

CN

Absolute stereochemistry. Rotation (+).

THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 22 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 10 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2007:1029651 CAPLUS

DN 147:365486

TI Preparation of 2-(phenylamino)thiazole derivatives as inhibitors of viral requaltum of 2-ipmemyamino) unamose derivetives as inhibit replication for the treatment of hepatitis C infection Zhang, Suoming; Phadke, Avinaen; Wang, Xiangshu; Liu, Cuixian Achillion Pharmaceuticals, Inc., USA PCT Int. Appl., 134pp. CODENT FIXED.

IN

PA

so

DT Patent

LA English

FAN.	TW	Τ.																
	PA:	TENT :	NO.			KIN	D	DATE			APPL	ICAT	ION	NO.		D.	ATE	
							-									-		
PI	WO	2007	1035	50		A2		2007	0913		WO 2	007-	US60	23		2	0070	308
	WO	2007	1035	50		A3		2007	1108									
		W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
			CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
			GE,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,
			KΡ,	KR,	KZ,	LA,	LC,	LK,	LR,	LS,	LT,	LU,	LY,	MA,	MD,	MG,	MK,	MN,
			MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RS,
			RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	SV,	SY,	TJ,	TM,	TN,	TR,	TT,	TZ,
			UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	zw							
		RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,
			IS,	IT,	LT,	LU,	LV,	MC,	MT,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,
								GA,										
			GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,
			BY,	KG,	KZ,	MD,	RU,	TJ,	TM,	AP,	EA,	EP,	OA					
	US	2007	0213	301		A1		2007	0913		US 2	007-	6837	49		2	0070	308
PRAI	US	2006	-780	609P		P		2006	0308									
os	MA	RPAT	147:	3654	B 6													
GI																		

AB Title compds. I (wherein A = (un)substituted Ph, benzyl, heteroxyl, etc.; R = CHO, CO(COOR, CO(COOR), etc.; R1 = (un)substituted haloslyy, haloslowy, alkylamino, etc.; R2 (0-2 substituents) = halo, OH, amino, etc.; R3 = H, halo, OH, etc.] and pharmaceutically acceptable salts thereof were prepared as inhibitors of viral replication. For instance, cyclocondensation of 3-bromosactylyytidine with N(4-pentable salts that the construction of the construction of

II 640281-90-9, Valopicitabine Ri: TRU (Therapeutic use), BIOL (Biological study); USES (Uses) (co-drug; preparation of (phenylamino)thiazoles as inhibitors of viral replication for treatment of hepatitis C infection.

RN 640281-90-9 CAPJUS CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

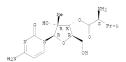
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111 ANSWER 11 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
AN
     2007:618644 CAPLUS
DN
     147:31277
     Polycyclic phenolic compounds and use in treating viral infections
IN
     Dugourd, Dominique
     Migenix Corporation, Can.
PA
so
     PCT Int. Appl., 77pp.
CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
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	PATENT	NO.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D.	ATE	
						-									-		
PI	WO 200	70625	28		A1		2007	0607		WO 2	006-	CA19	65		2	0061	201
	W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN, CO, C				CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE, GH, GI				HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,
		GE, GH, GM KP, KR, KI				LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY,	MA,	MD,	MG,	MK,
		MN,	MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,
		MN, MW, MX RS, RU, SC				SE,	SG,	SK,	SL,	SM,	SV,	SY,	TJ,	TM,	TN,	TR,	TT,
		TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	ZW						
	RW	: AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,

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IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, ST, SK, TR, BE, BJ, CR, CG, CI, CK, GA, GN, GQ, GW, ML, MR, ME, SN, TD, FG, BW, GH, KE, LS, MM, MD, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, WS 20070161611 Al 20070712 US 2006-565621 20061130 US 2006-565621 20061130 US 2006-565621 A 20061130 US 2006-565621 A 20061130 US 2006-565621 A 20061130 US 2006-565621 A 20061130
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- AB The present invention provides antiviral polycyclic phenolic compds.
 (PPCs) of formula I [R1 = R, skyl, aryl, cyclealky1, etc., R2 = R, OH, acyl, oxc, = (substituted) NH, SH, etc.] for use in treating or preventing viral infections and associated conditions, such as infections by Flaviviridae, Hapadneviridae, Respectificae, Papillomaviridae anoviridae, Orthomycoviridae, Permycoviridae, Permycoviridae, Permycoviridae, and Coronaviridae). Thus, II was prepared from estrone and 1-adamantanol, and inhibited viral release by 69% in BVDV-infected MDSR cells.
- IT 640281-90-9, Valopicitabine
 RL: THU (Therapeuric use); BIOL (Biological study); USES (Uses)
 (co-drug; estrone derive. for treatment of viral infections)
- RN 640281-90-9 CAPLUS
 CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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111 ANSWER 12 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
AN
     2007:412679
                  CAPLUS
DN
     146:395250
     Cyclosporin derivatives for the treatment and prevention of
     hepatitis C infection
     Houck, David Renwick
     Scynexis, Inc., USA
PCT Int. Appl., 65pp.
CODEN: PIXXD2
PA
so
     Patent
LA
    English
FAN.CNT 1
     PATENT NO.
                          KIND DATE
                                             APPLICATION NO.
                                                                       DATE
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PI	WO	2007	0416	32		A2			0412		WO 2	006-	US38	323		21	0061	002
	WO	2007				A3			1213									
		W:	ΑE,															
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
			GE,	SE, GH, G		HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,	KP,

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KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN,
              MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS,
              RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VC, VN, ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB,
                                                                     GR. HIL.
             IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
              GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AP, EA, EP, CA
     AU 2006299426
                           A1
                                  20070412
                                               AU 2006-299426
     US 20070173440
                           A1
                                  20070726
                                               US 2006-542930
                                                                       20061002
PRAI US 2005-722679P
                           P
     US 2006-787549P
                           P
                                  20060329
     WO 2006-US38823
                                  20061002
     MARPAT 146:395250
     This invention relates to 3-ether or 3-thioether derivs. of cyclosporin or
AB
     a pharmaceutically acceptable salt or solvate thereof, in combination with
     a second therapeutic agent for sequential or simultaneous administration
     in treatment and prevention of hepatitis C viral (
     HCV) infection. The second therapeutic agent is selected from
     modulators of NS3-4A protesse, modulators of NS5B RNA-dependent RNA
     polymerase, and immunomodulatory agents. Thus, treatment of
      1,4-diacetyl-3-methoxy-4-(gamma-hydroxymethylleucine)cyclosporin (275 mg)
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in methanol with 25 weight's sodium methoxide in methanol at room temperature yielded 33 mg of 3-methoxy-4-(gamma-hydroxymethylleucine)cyclosporin

The Compound T potently inhibited HCV replication in human liver cells to a greater extent than cyclosporin used as a control. In addition, when considering the level of cytotoxicity, the compound exhibited a wider safety margin (for example, cytotoxicity IC50 vs. antiviral EC50) than cyclosporine. The combination of Compound T and interferon-a was

additive. 640281-90-9, Valopicitabine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cyclosporin derivs, and their combinations for treatment and prevention of hepatitis C infection)

RN

(Compound I).

640281-90-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

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ANSWER 13 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
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AN 2007:320324 CAPLUS DN 146:394149

Valopicitabine dihydrochloride: a specific polymerase inhibitor of hepatitis C virus

Toniutto, Pierluigi; Fabris, Carlo; Bitetto, Davide; Fornasiere, Ezio;

Rapetti, Rachele; Firisi, Mario Internal Medicine, Medical Liver Transplantation Unit, DPMSC, University of Udine, Udine, 33100, Italy

90 Current Opinion in Investigational Drugs (Thomson Scientific) (2007), 8(2), 150-158

CODEN: COIDAZ; ISSN: 1472-4472

PB Thomson Scientific Journal; General Review

T.A English

A review. Idenix Pharmaceuticals Inc and Novartis AG are codeveloping valopicitabine dihydrochloride, a once-daily oral nucleoside for the potential treatment of BCV infection. In Jan. 2005, a phase IIa clin. trial comparing valopicitabine dihydrochloride with pegylated IFN in

treatment-naive HCV patients was ongoing, in addition to a phase IIb trial in patients that had previously failed pegylated IFN and ribavirin combination therapy. In Jan. 2006, an international phase III trial in treatment-refractory patients was planned for the first half of the year, with a phase III trial in treatment-naive individuals planned for the second half of the year.

640725-71-9, Valopicitabine dihydrochloride

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (valopicitabine dihydrochloride and interferon or ribavirin combination therapy was used to treat patient with hepatitis C virus infection)

640725-71-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HCl

RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 14 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2007:200728 CAPLUS 146:274570

Preparation of 2'-C-Me nucleoside 5 '-monophosphate and 4'-C-Me nucleoside 5'- monophosphate prodrugs for the treatment of hepatitis C viral infection

TN Erion, Mark D.; Reddy, K. Raja; Maccoss, Malcolm; Olsen, David B.

Merck & Co., Inc., USA; Metabasis Therapeutics, Inc. PCT Int. Appl., 268pp. CODEN: PIXXD2

Patent

FAN.		1 TENT	NO.			KIN		DATE			APPL					D.	ATE	
PI		2007				A2		2007			WO 2					2	0060	814
			CN, CO, C GE, GH, G KR, KZ, L MW, MX, M SC, SD, S US, UZ, V W: AT, BE, B IS, IT, L			AM, CU, HN, LC, NA, SG,	AT, CZ, HR, LK, NG, SK,	DE, HU, LR, NI, SL,	DK, ID, LS, NO, SM,	DM, IL, LT, NZ,	DZ, IN, LU, OM,	EC, IS, LV, PG,	EE, JP, LY, PH,	EG, KE, MA, PL,	ES, KG, MD, PT,	FI, KM, MG, RO,	GB, KN, MK, RS,	GD, KP, MN, RU,
		RW:	IS, CF, GM,	CG, KE,	LT, CI, LS,	LU, CM, MW,	LV, GA, MZ,	MC, GN, NA,	NL, GQ, SD,	PL, GW, SL,	PT, ML, SZ,	RO, MR, TZ,	SE, NE,	SI, SN,	SK, TD,	TR,	BF,	BJ, GH,
	CA	2618 1915	KG, KZ, MD 279720			A1 A2 CH, LT,	CY,	2007 2007 2008 CZ,	0222 0222 0430 DE,	DK,	AU 2 CA 2 EP 2 EE,	006- 006- 006- ES,	2618 8014 FI,	713 10 FR,	GΒ,	2 GR,	0060 0060 HU,	814 814 IE,
PRAI	US	2005						2005	0812									

US 2006-772649P 20060213 WO 2006-US31614 W 20060814 MARPAT 146:274570

AB 2'-C-Me nucleoside 5'-monophosphate and 4'-C-Me nucleoside 5'monophosphate prodrugs I, wherein B can be heterocyclic or heteroaryl rings; V is an optionally substituted monocyclic aryl or heteroaryl ring; W and W' are independently (un) substituted monocyclic aryl or heteroaryl rings, alkyl, aryl , heterocycloaryl or aralkyl groups; Ž is halo, cyano, keto, amido, etc. are prepared Further, I can also be prepared such that V and Z are connected via 3-5 atoms to form a cyclic group fused to aryl groups; Z and W connected via 3-5 atoms to form a cyclic group containing one heteroarom; or W and W' connected via an addnl. 2-5 atoms to form a cyclic group optionally containing 0-2 heteroatoms. Thus, II was prepared and tested for its in vitro activation in human liver microsomes by product capture (0.044 nmol/mg/min at activation 250 μM). I were also tested for their NTP accumulation in hepatocytes; in HCV-infected human liver assays; for tissue distribution following oral administration and the oral bioavailability in normal male rats.

926655-64-3P 926655-66-5P 926655-67-6P 926655-68-7P 926655-69-8P 926655-73-4P 926655-74-5P 926655-75-6P 926655-76-7P 926655-77-8P 926655-78-9P 926655-80-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(preparation of 2'-C-Me nucleoside 5'-monophosphate and 4'-C-Me nucleoside 5'- monophosphate prodrugs for the treatment of hepatitis C viral infection)

RN

926655-64-3 CAPLUS
Cytidine, 5'-0-[(2R, 48)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl)-2'-0-methyl-, 3'-octadecanoate (CA INDEX NAME)

926655-66-5 CAPLUS L-Valine, 3'-ester with 5'-0-[(2R,4S)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphocphorinan-2-yl]-2'-C-methylcytidine, 2,2,2-trifluoroacetate (1:?) (CA INDEX NAME)

CM

CRN 926655-65-4 CMF C24 H32 C1 N4 O9 P

Absolute stereochemistry.

CM

CRN 76-05-1 CMF C2 H F3 O2

926655-67-6 CAPLUS Cytidine, 5'-0-[(2R,48)-4-(3-chloropheny1)-2-oxido-1,3,2-dioxaphosphorinan-2-y1)-2'-0-methyl-, 3'-(2-methyl)-propanoate) (CA INDEX NAME)

926655-68-7 CAPLUS Cytidine, 5'-0-[(2R,45)-4-(3,5-difluorophenyl)-2-oxido-1,3,2dioxaphosphorinan-2-yl]-2'-C-methyl-, 3'-(2-ethylbutanoate) (CA INDEX MAMES

Absolute stereochemistry.

926655-69-8 CAPLUS Cytidine, 5'-0-6'(2R,4S)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl)-2'-0-methyl-,2',3'-bis(2-methylpropanoate) (CA INDEX NAME)

Absolute stereochemistry.

RN

szcob3-/3-4 CAPLUS
Cytidine, 5'-0-[(2R,4S)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl]-2'-C-methyl-N-[(pentyloxy)carbonyl]-, 3'-(2-methylpropanoate) (CA
INDEX NAME)

Absolute stereochemistry.

RN

926655-74-5 CAPLUS Cytidine, 5'-0-((2R, 4S)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl]-2'-0-methyl-N-((pentyloxy)carbonyl]-, 3'-butanoate (CA INDEX NAME) CAT

926655-75-6 CAPLUS Cytidine, 5'-0-(4-(3,5-difluorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl)-2'-C-nethyl-N-f (pentyloxy)carbonyl]-, 3'-(2-ethylbutanoate) (CA INDEX

Absolute stereochemistry.

RN

926655-76-7 CAPLUS Cyridine, 5'-0' (2R,48)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yi)-2'-C-methyl-N-E((pentyloxy)oarbonyl)-, 3'-acetate (CA INDEX NAME)

Absolute stereochemistry.

RN

926655-77-8 CAPLUS
Cytidine, 5'-0-[(2R,48)-4-(3-chlorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl)-2'-C-methyl-N-[(pentyloxy)carbonyl]-, 2',3'-diacetate (CA INDEX NAME)

Absolute stereochemistry.

926655-78-9 CAPUIS Cytidine, 5'-0-(4-(3,5-difluorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl)-2'-C-methyl-N-((pentyloxy)oarbonyl)-, 3'-acetate (CA INDEX NAME)

Absolute stereochemistry.

RN 926655-80-3 CAPLUS

N Cytidine, 5'-0-[4-(3,5-difluorophenyl)-2-oxido-1,3,2-dioxaphosphorinan-2-yl]-2'-C-methyl-N-[(pentyloxy|carbonyl]-, 2',3'-diacetate (CA INDEX NAME)

Absolute stereochemistry.

2007:85871 CAPLUS

146:177157

CA 2615626

EP 1909564

2.

PRAI US 2005-700475P

US 2006-776640P

WO 2006-US27485

AN

DN

Small animal model for HCV replication Weiner, Amy; Aukerman, Sharon Lea; Mendel, Dirk; Zhu, Qing

111 ANSWER 15 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

Al

A2

-

P

W

An animal model for HCV (hepatitis C virus)

BA, HR, MK, RS

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PA
     Novartis A.-G., Switz.
     PCT Int. Appl., 85pp.
     CODEN: PIXXD2
DТ
     Patent
LA
     English
FAN.CNT
     PATENT NO.
                           KIND
                                  DATE
                                               APPLICATION NO.
                                                                         DATE
ΡI
     WO 2007011777
                            A2
                                               WO 2006-US27485
                                                                         20060715
     WO 2007011777
                            A3
         W: AE, AG, AL,
                          AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
              CN, CO, CR,
                           CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
              GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP,
              KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN,
              MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG,
              US, UZ,
                           VN, ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY,
                                   CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
              IS, IT,
                               LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
                      LT, LU,
              CF, CG, CI,
                           CM,
                               GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
              GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU,
                              TJ, TM, AP, EA, EP, OA
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20080416

20060223

20060715

CA 2006-2615626 EP 2006-787397

AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL,

replication and/or production of virus or virus like particles is provided. The invention utilizes an HCV replicon present in a cell to

20060715

20060715

AB

deliver HCV nucleic acid and replicate and express HCV proteins in an animal model comprising an animal that has been immunocompromised. The invention further provides a method of treatment or prevention of HCV in a mammal which comprises administering to the mammal a combination which comprises an immunomodulatory compound and another antiviral agent. Also provided are cell lines showing a decreased sensitivity to interferon alpha or some other immunomodulator and methods of making or isolating such cell lines. 640281-90-9, Valopicitabine RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(small animal model for hepatitis C virus

replication)

640281-90-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

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L11 ANSWER 16 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
ΔN
    2006:1283521 CAPLUS
DN
    146:20343
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P

Use of 4-[3,5-Bis-(2-hydroxyphenyl)-[1,2,4]-triazol-l-yl]benzoic acid for the treatment of liver diseases in which iron plays a role in pathogenesis Alberti, Daniele; Marks, Peter; Nick, Hanspeter; Rojkjaer, Lisa Grace TN

PA Novartis AG, Switz.; Novartis Pharma GmbH so PCT Int. Appl., 20pp. CODEN: PIXXD2

DТ Patent

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PI	WO	2006	1305	32		A2		2006	1207			2006-				2	0060	530
	WO	2006	1305	32		A3		2007	1122									
		W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	, BG,	BR,	BW,	BY,	BZ,	CA,	CH,
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	AU	2006	2527	18		A1		2006	1207		AU :	2006-	2527	18		2	0060	530
	CA	2608	709			A1		2006	1207		CA :	2006-	2608	709		2	0060	530
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		2005																
		2006						2006										

20060530

WO 2006-US20677

10/609.298

- The invention discloses use of 4-[3,5-Bis-(2-hydroxyphenyl)-[1,2,4]triazol-1-yl]benzoic acid for the manufacture of pharmaceutical compns. for the treatment of liver diseases in humans in which iron plays a role in pathogenesis, including viral diseases, e.g. chronic hepatitis C, optionally in conjunction with antiviral agents and for the treatment of nonviral diseases, e.g. non-alc. steatchepatitis and non-alc.
- fatty liver disease. 640281-90-9, Valopicitabine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(triazolyl benzoic acid derivative for treatment of liver diseases in which iron plays role in pathogenesis) 640281-90-9 CAPLUS - L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

- ANSWER 17 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2006:1257296 CAPLUS DN 146:54578
- Recent patents on nucleoside and nucleotide inhibitors for HCV
- Shim, Jae H.; Hong, Zhi; Wu, Jim Z. AU
- Drug Discovery, Valeant Pharmaceuticals International, Costa Mesa, CA, 92626. USA
- Recent Patents on Anti-Infective Drug Discovery (2006), 1(3), 323-331 CODEN: RPADCX; ISSN: 1574-891X
- DR Bentham Science Publishers Ltd.
- Journal; General Review
- English
- A review. Hepatitis C virus (HCV) infection 28
 - is a leading cause of liver diseases such as cirrhosis and hepatocellular There are estimated 170 million people worldwide chronically infected with the virus. The lack of highly effective and safe therapeutics for HCV infection has spurred intensive efforts to develop anti-HCV drugs as evidenced by the large number of new patent applications filed each year. Nucleoside and nucleotide inhibitors are the analogs of DNA or RNA substrates, and they inhibit viral polymerases by acting as chain terminators, viral mutagens, or simple competitive inhibitors. The successful development of various nucleoside and nucleotide inhibitors for the treatment of HIV and HBV infections has prompted the drug industry to seek similar strategies for HCV. This review summarizes recently issued or published patents covering nucleoside and nucleotide inhibitors for HCV. The claimed chemical structures and available biol. activities, mechanism of action, and drug resistance profiles are discussed. The development status of several promising nucleoside inhibitors is also described.
- 640725-71-9, NM283
 - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (recent patents on nucleoside and nucleotide inhibitors for HCV
- 640725-71-9 CAPLUS
- L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HC1

RE.CNT 45 THERE ARE 45 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 18 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2006:1086375 CAPLUS

DN 146:54739

Ribavirin antagonizes the in vitro anti-hepatitis C

virus activity of 2'-C-methylcytidine, the active component of valopicitabine

Coelmont, Lotte; Paeshuyse, Jan; Windisch, Marc P.; De Clercq, Erik; Bartenschlager, Ralf; Neyts, Johan

Rega Institute for Medical Research, KULeuven, Louvain, 3000, Belg. SO Antimicrobial Agents and Chemotherapy (2006), 50(10), 3444-3446

CODEN: AMACCO; ISSN: 0066-4804 PB American Society for Microbiology

DT Journal

LA English

Ribavirin antagonizes the in vitro anti-hepatitis C

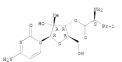
virus (HCV) activity of the pyrimidine nucleoside analog 2'-C-methylcytidine, the active component of the exptl. anti-HCV drug valopicitabine. In contrast, the combination of ribavirin with either the purine nucleoside analog 2'-C-methyladenosine or the HCV protease inhibitor VX-950 resulted in an additive antiviral activity. These findings may have implications when planning clin. studies with valopicitabine.

640281-90-9D, Valopicitabine, metabolite RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ribavirin antagonizes anti-hepatitis C virus

activity of 2'-C-methylcytidine, active component of valopicitabine) 640281-90-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 13 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 19 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

2006:1045856 CAPLUS AN

DN 146:28022

Synthesis and Pharmacokinetics of Valopicitabine (NM283), an Efficient

Prodrug of the Potent Anti-HCV Agent 2'-C-Methylcytidine Pierra, Claire; Amador, Agnes; Benzaria, Samira; Cretton-Scott, Erika; D'Amours, Marc; Mao, John; Mathieu, Steven; Moussa, Adel; Bridges, Edward

- G.; Standring, David N.; Sommadossi, Jean-Pierre; Storer, Richard; Gosselin, Gilles
- CS Laboratoire Cooperatif Idenix-CNRS-Universite Montpellier II Case Courrier 008, Universite Montpellier II, Montpellier, 34095, Fr. SO Journal of Medicinal Chemistry (2006), 49(22), 6614-6620
- CODEN: JMCMAR; ISSN: 0022-2623
- PB American Chemical Society
- DT Journal LA English
- OS CASREACT 146:28022
- AB In the search for new therapeutic agents against chronic hepatitis C, 2'-C-methyloytidine was discovered to be a potent and selective inhibitor in cell culture of a number of RNA viruses, including the

pestivirus bovine viral diarrhea virus, a surrogate model for hepatitis C virus (HCV), and three flaviviruses, namely, yellow fever virus, West Nile virus, and

riavivaruses, namely, yellow rever virus, west kile virus, and dengue-2 virus. However, pharmacokinetic studies revealed that 2'-C-methylcytidine suffers from a low oral bicavailability. To overcome this limitation, the authors have synthesized the 3'-O-L-valinyl ester derivative (NM-283; dihydrochloride salt of valopicitabine) of

2'-C-methyl-cytidine. The authors present the chemical synthesis and physicochem. characteristics of NM-183, anti-HCV prodrug candidate, as well as a comparative study of its pharmacokinetic parameters with those of its parent nucleoside analog, 2'-C-methyl-cytidine.

IT 640725-71-9P

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); BTO. (Siological study); PREP (Preparation) (preparation and pharmacokinetics of NM-283, a prodrug of anti-MCV agent 2'-Co-methyleytidine)

RN 640725-71-9 CAPLUS CN L-Valine, 3 ester with 2 -C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HCl

IT 640725-70-8P 642075-44-3P

RL: RCI (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACI (Reactant or reagent) (preparation and pharmacokinetics of NM-283, a prodrug of anti-HCV

agent 2'-C-methylcytidine) 640725-70-8 CAPLUS

CN L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

642075-44-3 CAPLUS RN

L-Valine, N-[(1,1-dimethylethoxy)oarbonyl]-, 3'-ester with 5'-O-[(1,1-dimethylethyl)diphenylsilyl]-N-[(dimethylamino)methylene]-2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT RE.CNT 29

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L11 ANSWER 20 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2006\!:\!945608 CAPLUS
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DN 145:315225

TI

Bicyclic nucleosides and nucleotides as therapeutic agents Francom, Paula; Nearn, Roland Henry; Draffan, Alistair George; Lambert, John Nicholas; Bond, Silas

Biota, Inc., USA PCT Int. Appl., 107pp. CODEN: PIXXD2

so

DT Patent

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PI		2006				Al		2006									0060	
		W:	CN, GE, KZ, MZ, SG,	CO, GH, LC, NA, SK,	CR, GM, LK, NG, SL,	CU, HR, LR, NI,	CZ, HU, LS, NO, SY,	AU, DE, ID, LT, NZ, TJ,	DK, IL, LU, OM,	DM, IN, LV, PG,	DZ, IS, LY, PH,	EC, JP, MA, PL,	EE, KE, MD, PT,	EG, KG, MG, RO,	ES, KM, MK, RU,	FI, KN, MN, SC,	GB, KP, MW, SD,	GD, KR, MX, SE,
		RW:	IS, CF, GM,	CG, KE,	LT, CI, LS,	LU,	LV. GA. MZ.	CZ, MC, GN, NA, TM	NL, GQ,	PL, GW,	PT, ML,	RO, MR,	SE, NE,	SI, SN,	SK, TD,	TR, TG,	BF, BW,	BJ, GH,
	CA	2006 2600 1858	886			Al Al		2006 2006 2007	0914		CA 2	006 006 006-	2600	886		2	0060 0060	308

10/609.298

R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LIT, LU, LV, MC, NI, PL, PT, RO, SE, SI, SK, TR PRAI US 2005-6616659 P 2005038 WO 2006-AU303 T.F 20060308

CASREACT 145:315225

The invention relates to the use of bicyclic nucleosides and nucleotides I, wherein: A is O, S, CH2, CHF, CF2 or NR; B is bicyclic heterocycle; R1, R2, R2', R3, R3', and R4 are independently H, halogen, OH, N3, CN, alkyl, alkenyl, alkynyl, aryl, acyl, arylalkyl, heterocyclyl, heteroaryl, cycloalkyl, cycloalkenyl, alkyloxy, alkenyloxy, alkynoxy, aryloxy, acyloxy, oxyacyl, arylalkoxy, heterocycloxy, heteroaryloxy, cycloalkoxy, cycloalkenoxy, amino, aminoacyl, aminoacyloxy, acylamino, oxyacylamino, oxyacyloxy, acylimino, acyliminoxy, oxyacylimino, aminothioacyl, Oxyadylady, adylamino, aminosulfony, thio, thioacyla, thioacylanino, aminosulfony, thio, thioacyla, thioacylanino, axinosulfony, oxythioacyl, oxythioacylanino, oxythioacylani Y' is H, halogen, N3, Me, Et or CN; R4' is -CY2SH, -CY2OH, -CY2NH, or L'-R5; L' is selected from the group consisting of -CY2-, -CY2CY2-, -CY2CY2-, and -CY2NHCY2-; Y is H, OR, halogen, alkyl, alkenyl, alkynyl; R5 is OR, NR2, monophosphate, diphosphate, and triphosphate, or a mono, di or triphosphate mimic; each R is independently H, CF3, alkyl, alkenyl, alkynyl, aryl, acyl, cycloalkyl, cycloalkenyl, heteroaryl, heterocyclyl; were prepared for the treatment of infectious diseases, and in particular, viral infections. Title compds. were typically active in the replicon assay in the range 1 to >1000 μM and cytotoxic in the range 30 to >100 μM. HCV-polymerase inhibition by title compds. is

also reported. 23643-36-9P 909394-67-8P 909394-72-5P 909394-73-6P 909394-74-7P 909394-75-8P 909394-76-9P 909394-77-0P 909394-81-6P 909394-82-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of bicyclic nucleosides and nucleotides as therapeutic agents) 23643-36-9 CAPLUS

2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-benzoyl-2-C-methyl- β -Dribofuranosvl) - (9CI) (CA INDEX NAME)

RN 909394-67-8 CAPUUS CN Cytidine, 5-lode-2"-C-methyl-, 2",3",5"-tribenzoate (9CI) (CA INDEX NAME) Absolute stereochemistry.

RN 909394-72-5 CAPLUS CN Cytidine, N-methyl-2'-C-methyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 909394-73-6 CAPLUS CN Cytidine, 2'-C-methyl-N-(phenylmethyl)-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

RN 909394-74-7 CAPLUS CN Cytidine, 2'-C-methyl-N-propyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 909394-75-8 CAPLUS
CN Cytidine, S-iodo-N-methyl-2'-C-methyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 909394-76-9 CAPLUS CN Cytidine, 5-iodo-2'-C-methyl-N-(phenylmethyl)-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

RN 909394-77-0 CAPLUS CN Cytidine, 5-iodo-2'-C-methyl-N-propyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 909394-81-6 CAPLUS
CN Cytidine, 5-(3-hydroxy-1-propynyl)-2'-C-methyl-, 2',3',5'-tribenzoate
(9C1) (CA INDEX NAME)

Absolute stereochemistry.

RN 909394-82-7 CAPLUS CN Cytidine, 2'-C-methyl-5-(3-oxo-1-propynyl)-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT RE.CNT 28

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L11 ANSWER 21 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN -2006\!:\!894484 CAPLUS
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DN 145:285094 TI

Glucosidase inhibitor combinations with adjunctive therapies for treating or preventing Flaviviridae infections

IN Dugourd, Dominique; Rubinchik, Evelina; Clement, Jacob; Friedland, Hillel David

PA Migenix Inc., Can.

U.S. Pat. Appl. Publ., 69pp. CODEN: USXXCO SO

DT Patent

English LA

FAN.	CNT	1																
	PA1	TENT :	NO.			KIN	0	DATE			APF	LICAT	ION :	NO.		D.	ATE	
PI		2006										2006-						
	AU	2006	2210	80		A1		2006	0914		ΑU	2006-	2210	80		2	0060	209
	CA	2597	213			A1		2006	0914		CA	2006-	2597	213		2	0060	209
	WO	2006	0962	85		A2		2006	0914		WO	2006-	US49	27		2	0060	209
	WO	2006	0962	85		A3		2007	0125									
		W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BE	3, BG,	BR,	BW,	BY,	BZ,	CA,	CH,
			CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ	, EC,	EE,	EG,	ES,	FI,	GB,	GD,
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS	JP,	KE,	KG,	KM,	KN,	KP,	KR,
			KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY	, MA,	MD,	MG,	MK,	MN,	MW,	MX,
			MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH	, PL,	PT,	RO,	RU,	SC,	SD,	SE,
			SG,	SK,	SL,	SM,	SY,	ΤJ,	TM,	TN,	TF	, II,	TZ,	UA,	UG,	US,	UZ,	VC,
			VN,	YU,	ZA,	ZM,	ZW											
		RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE	, ES,	FI,	FR,	GB,	GR,	HU,	IE,
			IS,	IT,	LT,	LU,	LV,	MC,	NL,	PL,	PI	, RO,	SE,	SI,	SK,	TR,	BF,	BJ,
			CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	MI	, MR,	NE,	SN,	TD,	TG,	BW,	GH,
			GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ	, TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
						RU,												
	EP	1853	317			A2		2007	1114		ΕP	2006-	7482	02		2	0060	209
		R:										E, ES,						IE,
												, PT,						
												2007-						
	IN	2007	KN03	225		A		2008	0321		IN	2007-	KN32	25		2	0070	831
	KR	2007	1027	41		A		2007	1019		KR	2007-	7205	40		2	0070	907
PRAI	US	2005	-651	910P		P		2005	0209									
	US	2005	-664	297₽		₽		2005	0321									
	US	2005	-735	464P		P		2005	1112									
	WO	2006	-US4	927		W		2006	0209									

McIntosh

GI

The present disclosure relates generally to compns. having a glucosidase inhibitor [castanospermine (I) or a derivative thereof, such as celgosivir] in combination with adjunctive therapies of compds. that alter immune function (such as interferon) and compds. that alter viral replication (such as nucleoside analogs like ribavirin), which can be used to treat or prevent infections caused by or associated with a virus of the Flaviviridae family, particularly infections caused by or associated with Hepatitis C virus (HCV). Examples include synergy of castanospermine or celgosivir in combination with other drugs such as interferons in a checkboard approach.

640725-71-9, NM283 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (glucosidase inhibitor combinations with adjunctive therapies for

treating or preventing Flaviviridae infections) RN 640725-71-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HCl

- ANSWER 22 OF 37 CAPLUS COPYRIGHT 2008 ACS on SIN
- 2006:774481 CAPLUS AN
- DΝ 146:402230
- Synthesis of 2'-C-methylcytidine and 2'-C-methyluridine derivatives modified in the 3'-position as potential antiviral agents Plierra, Claire; Amador, Agnes; Badaroux, Eric; Storer, Richard; Gosselin,
- AU
- Laboratoire Cooperatif Idenix-CNRS-Universite Montpellier II, Universite Montpellier II, Montpellier, 34095/5, Fr.
 Collection of Czechoslovak Chemical Communications (2006), 71(7), 991-1010 so
- CODEN: CCCCAK; ISSN: 0010-0765 Institute of Organic Chemistry and Biochemistry, Academy of Sciences of PB the Czech Republic
 - Journal
- LA English
- os CASREACT 146:402230

2'-C-methylcytidine and 2'-C-methyluridine derivs. modified in the 3'-position, e.g. I-HCl, were prepared via Barton deoxygenation, amination, stereoselective cyclization, ring opening and fluorination from 2'-C-methyluridine or uridine. The antiviral activity of the title compds. was tested against RNA viruses and was found to be inactive. was found that the modification at the 3'-position resulted in loss of

10/609.298

antiviral activity.

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or readent)

(no activity; preparation and antiviral activity of methylcytidine and methyluridine derivs. via Barton deoxygenation, amination, stereoselective cyclization, ring opening and fluorination from methyluridine or uridine)

934014-31-0 CAPLUS Uridine, 2'-C-methyl-3'-O-methyl- (CA INDEX NAME) CN

Absolute stereochemistry.

934014-32-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (no activity; preparation and antiviral activity of methylcytidine and

methyluridine derivs. via Barton deoxygenation, amination, stereoselective cyclization, ring opening and fluorination from

methyluridine or uridine) 934014-32-1 CAPLUS

Cytidine, 2'-C-methyl-3'-O-methyl-, hydrochloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.

● HCl

934014-23-0P 934014-24-1P 934014-27-4P 934014-28-5P 934014-30-9P 934014-42-3P

RL: RCI (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACI

(Reactant or reagent) (preparation and antiviral activity of methylcytidine and methyluridine derivs. via Barton deoxygenation, amination, stereoselective cyclization, ring opening and fluorination from methyluridine or

uridine) 934014-23-0 CAPLUS DM

Uridine, 5'-0-[(1,1-dimethylethyl)dimethylsilyl]-2'-C-methyl-2'-0-(tetrahydro-2H-pyran-2-yl)-, 3'-methanesulfonate (CA INDEX NAME)

934014-24-1 CAPLUS Uridine, 5'-0-[(1,1-dimethylethyl)dimethylsilyl]-2'-C-methyl-, 3'-methanesulfonate (CA INDEX NAME)

Absolute stereochemistry.

934014-27-4 CAPLUS 2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-acetyl-2-C-methyl- β -D-RN CN xylofuranosyl) - (CA INDEX NAME)

Absolute stereochemistry.

934014-28-5 CAPLUS 2(1H)-Pyrimidinos, 3,4-dihydro-4-thioxo-1-(2,3,5-tri-O-acetyl-2-C-methyl-#-D-xylofuranosyl)- (CA INDEX NAME)

Absolute stereochemistry.

934014-30-9 CAPLUS

Uridine, 5'-0-[(1,1-dimethylethyl)dimethylsilyl]-2'-C-methyl-3'-O-methyl-2'-O-(tetrahydro-2H-pyran-2-yl)- (CA INDEX NAME)

10/609.298

Absolute stereochemistry.

934014-42-3 CAPLUS

CN Cvtidine, 2'-C-methyl-3'-O-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 23 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2006:708636 CAPLUS

DN 146:19039

Valopicitabine: anti-hepatitis C virus drug RNA-directed RNA polymerase (NS5B) inhibitor

Sorbera, L. A.; Castaner, J.; Leeson, P. A. Prous Science, Barcelona, 08080, Spain

CS

Drugs of the Future (2006), 31(4), 320-324 CODEN: DRFUD4; ISSN: 0377-8282

Prous Science

Journal: General Review

īΑ English

AB A review. Chronic hepatitis C is caused by infection

with the hepatitis C virus (HCV), a member of the Flaviviridae family of viruses. Currently available

treatment for HCV, including the standard combination therapy with interferon and ribavirin, is often unsuccessful at eradicating infection. In addition, the therapies now used to treat chronic hepatitis

C are associated with substantial side effects. Therefore, new therapeutic strategies such as the use of antiviral drugs targeted to HCV-specific viral enzymes are being explored. One such option is the RNA-directed RNA polymerase (NSSB) inhibitor valopicitabine (NM-283). an orally bicavailable prodrug of the novel ribonucleoside analog NM-107.

This compound has shown in vitro activity against HCV-related bovine viral diarrhea virus (BVDV) polymerase. In patients with HCV-1 infection, valopicitabine produced redns. in HCV RNA viral load when administered either as monotherapy or in combination with pegylated interferon. When used together, valopicitabine and interferon appear to have synergistic antiviral effects both in vitro and in vivo. The compound is generally well tolerated, with gastrointestinal

effects being the most commonly observed treatment-related adverse events. 640725-71-9P, NM-283 RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU

(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(valopicitabine reduced HCV RNA viral load either as

monotherapy or in combination with pegylated interferon against HCV-related bovine viral diarrhea virus polymerase and in patient with hepatitis C virus-1 infection)

640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HC1

RE.CNT 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 24 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2006:425398 CAPLUS DN 145:39734

Nucleoside analog inhibitors of hepatitis C virus

replication

Carroll, S. S.; Olsen, D. B.

Department of Antiviral Research, Merck Research Laboratories, West Point, PA, 19486, USA

so Infectious Disorders: Drug Targets (2006), 6(1), 17-29 CODEN: IDDTAD; ISSN: 1871-5265

PB Bentham Science Publishers Ltd.

Journal; General Review

LA

English A review. Of the 30 compds, currently marketed in the United States for treatment of viral infections, 15 are nucleoside analogs, demonstrating the utility of this class of compound as a source of antiviral drugs. The success of nucleoside analogs in treating other viral infections provides a compelling rationale for the significant effort that is currently being devoted to the discovery and development of nucleoside analogs to treat infection by hepatitis C virus (HCV) that

may lead to improvements in response rates compared to currently available therapies. Several different approaches were adopted to identify promising analogs, including the use of surrogate viruses in cell culture assays, screening in the cell-based bicistronic HCV replicon assay, and screening nucleoside triphosphates for the ability to inhibit the activity of the HCV RNA-dependent RNA polymerase in vitro.

Several classes of ribonucleoside analogs with modifications of the ribose inhibit HCV replication. Nucleoside analogs incorporating a 2'-C-Me modification are potent inhibitors in the replicon assay in the

absence of cytotoxicity, and appear to exert their inhibition by acting as functional chain terminators of RNA synthesis. NM283, a prodrug of 2'-C-methylcytidine, has entered clin. trials and demonstrated viral load redns. in subjects infected with genotype 1 HCV, a genotype known to be difficult to treat effectively with currently approved

therapies. Overall, results to date offer encouragement that improved therapies to treat HCV infection including newly developed nucleoside analogs may become available within the next few years.

640725-71-9, NM 283 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nucleoside analog inhibitors of hepatitis C virus

replication; 640725-71-9 CAPLUS RN

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

●2 HC1

THERE ARE 65 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 65 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 25 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2006:342840 CAPLUS

DN 144:381956

Combination antiviral compositions comprising castanospermine and use for the treatment and prevention of infections caused by or associated with a virus of the Flaviviridae family Dugourd, Dominique Migenix Inc., Can.

IN PA

so PCT Int. Appl., 54 pp. CODEN: PIXXD2

DT Patent

		TENT 1															
		2006														0051	
PI	WO																
		w:					AU,										
							DE,										
							ID,										
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							MC,										
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							2006										
		1802	1093	5//	Al		2006	0504		US 2	005-	2448	11		2	0051	
	EP						2007										
		R:					CZ,										
							LV,										
	CN	1010	3333.		A		2007	0912		CN 2	005-	8003	4258		2		
	JP	2008	0158.	16	T		2008	0515		JP 2	007-	5349	81		2	0051	006
	MX	2007	3385.	3	A		2007	1121		MX 2	007-	3853			- 2	0070	329
	KR	2007	1618	/9	A		2007	0614		KR 2	007-	7087	15		2	0070	417
	TM	20071	KNUI.	333	A		2007	0720		IN 2	007-	KN13	5.5		- 2	0070	41/
PKAI	US	2004-	-6T6	1815	2		2004	T006									
AB																	
		inv															

amounter therapeutic agent to treat or prevent inhections based a associated with a virus of the Flaviviridae family, particularly infections caused by or associated with Hepatitis C virus (RCV), and to the use of such compds. to examine the biol. mechanisms of HCV infection.

882489-96-5

RL: BSU (Biological study, unclassified); BIOL (Biological study) (castanospermine-containing combination antiviral compns., and use for treatment of Flaviviridae infections)

882489-96-5 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, mixt. with (1S,6S,7R,8R,8aR)-octahydro-1,6,7,8-indolizinetetrol (9CI) (CA INDEX NAME 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 79831-76-8 CMF C8 H15 N O4

Absolute stereochemistry. Rotation (+).

640281-90-9, Valopicitabine

RE: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(castanospermine-containing combination antiviral compns., and use for treatment of Flaviviridae infections)
640281-90-9 CAPLUS
L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 4 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 26 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN 2006:149315 CAPLUS AN

DN 144:205728

Methods using a Type II interferon receptor agonist alone or in combination with a direct antiviral drug for treating hepatitis C virus infection

Blatt, Lawrence M.

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PA
        Intermune, Inc., USA
PCT Int. Appl., 139 pp.
CODEN: PIXXD2
         Datent
T.A
        English
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	CNT 1																
	PATENT	NO.			KIN	D	DATE			APPL	ICAT	ION	NO.		D.	ATE	
						-									-		
PI	WO 2006	0169	30		A2		2006	0216	1	WO 2	005-	US16	927		2	0050	513
	WO 2006	0169	30		A3		2006	0803									
	W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KP,	KR,	KZ,
		LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA.
		NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	sc,	SD,	SE,	SG,	SK,
		SL,	SM,	SY,	TJ,	TM,	TN,	TR.	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,
		ZA,	ZM,	ZW													
	RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,
		IS,	IT,	LT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	BJ,	CF,
		CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG,	BW,	GH,	GM,
		KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,	KG,
		KZ,	MD,	RU,	TJ,	TM											

PRAI US 2004-571322P AB

P 20040514 The invention provides methods for treating hepatitis C virus (HCV) infection; methods for reducing the incidence of complications associated with HCV and cirrhosis of the liver; and methods for reducing viral load, or reducing the time to viral clearance, or reducing morbidity or mortality in the clin. outcomes, in patients suffering from HCV infection. The methods generally involve administering to the individual a Type II interferon receptor agonist alone or in combination with a direct antiviral drug.

640725-71-9, NM 283 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Elological study); USES (Uses)
(type II interferon receptor agonist alone or in combination with direct antiviral drug for treating hepatitis C

virus infection) 640725-71-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

● 2 HCl

111 ANSWER 27 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2006:103884 CAPLUS

DN 144:171198

Preparation of alkyl-substituted 2-deoxy-2-fluoro-D-ribofuranosyl TI pyrimidine and purine nucleoside analogs via condensation of the lactone

Wang, Peiyuan; Stec, Wojciech; Clark, Jeremy; Chun, Byoung-Kwon; Shi, IN Junxing; Du, Jinfa Pharmasset, Inc., USA

PA

PCT Int. Appl., 34 pp. CODEN: PIXXD2

Datent

LA English

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		CENT :				KIN		DATE									ATE	
DT		2006										2005					0050	
FI		2006									WO 2	-005-	0020	210		-	0050	121
											BB.	BG.	BR.	BW.	BY.	BZ.	CA.	CH,
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			KG,	KΖ,	MD,	RU,	IJ,	TM										
	AU	2005	2670	51		Al		2006	0202		AU 2	2005-	2670	51		- 2	0050	721
		2574				AL		2006	0202		CA 2	2005-	20/4	62T			0050	
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		20.										PI.						
	CN	1010 2005	2309	4	,	A		2007	0822	,	CN 2	2005-	8003	1530	,	2	0050	721
	BR	2005	0121	0.4		A		2008	0311		BR 2	2005-	1210	4		2	0050	721
	JP	2008	5075	47		T		2008	0313		JP 2	2007-	5227	63		2	0050	721
	US	2008 2006 2007 2007	0199	783		Al		2006	0907		US 2	2006-	3535	97		2	0060	213
	MX	2007	0080	3		A		2007	0402		MX 2	2007-	803			2	0070	119
	IN	2007	KNOU	605		A		2007	1202		IN 2	2007-	KN60	5		2	0070	220
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		2005																
		2005				W		2005	0721									
os	MA	RPAT	144:	1711	98													
GI																		

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

A process for preparing of 2-deoxy-2-fluoro-2-methyl-D-ribonolactones, I, wherein R1 and R2 can independently be H, CH3, acetyl, benzoyl, pivaloyl, 4-nitrobenzoyl, 3-nitrobenzoyl, 2-nitrobenzoyl, 4-chlorobenzoyl, 3-chlorobenzoyl, 2-chlorobenzoyl, 4-methylbenzoyl, 3-methylbenzoyl, 2-methylbenzoyl, 4-phenylbenzoyl, benzyl, 4-methoxybenzyl, trityl, trialkylsilyl, t-butyl-dialkylsilyl, t-butyl-dialkylsilyl, t-butyl-dialkylsilyl, t-butyl-diphenylsilyl, TIPDS, THP, MOM, or MEM are prepared and used in the condensation to 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs. Thus, 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs II and III, wherein X is a halogen; Y is N or CH; Z is a halogen, hydroxyl, ether, thiol, thioether, (un)substituted amine or alkyl; Rl' is alkyl, vinyl, ethynyl; Rl' and R3' can be same or different H, alkyl, arylalkyl, acyl, cyclic acetal such as 2',3'-O-isopropylidene or 2',3-0-benzylidene, or 2',3'-cyclic carbonate; R4, R5, and R6 are independently H, halogen, hydroxyl, ether, thiol, thioether, N3, (un) substituted amine, (un) substituted amido, alkyl, halogenated alkyl, alkenyl, halogenated alkenyl, alkynyl, halogenated alkynyl, hydroxy alkyl, alkoxy are prepared and are potential anti-HCV agents. Specifically, IV was prepared (no yield, claimed) via condensation, alkylation and stereoselective fluorination reactions and can exhibit potential use as an anti-HCV agent. 874638-81-0P

If S/4638-81-UP (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RADT (Reactant or reagent) (preparation of a RMJ-substituted 2-deoxy-2-fluoro-D-ribofuranosyl pyrimidine and purine nucleoside analogs via condensation of the lactone to nucleoside)

RN 874638-81-0 CAPLUS

Benzamide, N-[1-[5-0-benzoyl-2-C-methyl-3-0-(methylsulfonyl)-β-Darabinofuranosyl]-1,2-dihydro-2-oxo-4-pyrimidinyl]- (CA INDEX NAME)

Absolute stereochemistry.

- ANSWER 28 OF 37 CAPLUS COPYRIGHT 2008 ACS on SIN
- AN 2005:1151389 CAPLUS 145:271979
- DN
- NM 283, an efficient prodrug of the potent anti-HCV agent 2'-C-methylcytidine
- Pierra, C.; Benzaria, S.; Amador, A.; Moussa, A.; Mathieu, S.; Storer, R.; Gosselin, G.
 - Laboratoire Cooperatif Idenix, CNRS, Universite Montpellier II,
- Montpellier, 5, Fr.
 Nucleosides, Nucleotides & Nucleic Acids (2005), 24(5-7), 767-770 so
- CODEN: NNNAFY; ISSN: 1525-7770 Taylor & Francis, Inc. PB
- Journal
- English
- CASREACT 145:271979
- AB In order to improve the oral bioavailability of 2'-C-methylcytidine, a
- potent anti-ECV agent, the corresponding 3*-O-L-valinyl ester derivative (MN 233) has been synthesized. Based on its ease of synthesis and its physicochem. properties, NN 263 has emerged as a promising antiviral drug for treatment of chronic MCV infection. 23643-36-96 467075-70-89
- RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
- (Reactant or reagent) (preparation of NM 283 as efficient prodrug of potent anti-HCV
- agent 2'-C-methylcytidine)
- 23643-36-9 CAPLUS 2.4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-0-benzoyl-2-C-methyl- β -D-ribofuranosyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- 640725-70-8 CAPLUS DAT
- L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

640725-71-9P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prodrug; preparation of NM 283 as efficient prodrug of potent anti-HCV agent 2'-C-methylcytidine)

640725-71-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HC1

RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 29 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:684531 CAPLUS

DΝ 143:431740

Emerging drugs for chronic hepatitis C

Bhopale, Girish Mahadeorao; Nanda, Rabindra Kumar Research and Development Division, Hindustan Antibiotics Limited, Pimpri. CS

Pune, 411018, India so Hepatology Research (2005), 32(3), 146-153

CODEN: HPRSFM; ISSN: 1386-6346

PB Elsevier B.V.

Journal; General Review

English LA

A review. Hepatitis C virus (HCV) is a

major cause of chronic hepatitis, liver cirrhosis and hepatocellular carcinoma worldwide. A combination therapy comprising pegylated interferon and ribavirin currently represents the most effective therapy for chronic HCV infection. The limitations of this current therapy mainly its efficacy and significant side effects have prompted the development of new drugs. Few categories of therapeutic agents appear promising for future therapy, e.g. novel interferons, ribavirin analogs, antisense oligonucleotides, short interfering RNAs, ribozymes, enzyme inhibitors, immunomodulatory agents, antifibrotic agents, therapeutic vaccines and antibodies. Few drugs belong to afore-mentioned categories have already reached the different clin. phases of development. The present article highlights the status of current available therapies and emerging drugs for the treatment of hepatitis C. 640725-71-9, NM 283

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses) (NM283 proved promising therapeutic effect in treating chronic hepatitis C patient)

640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

2 HC1

RE.CNT 64 THERE ARE 64 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

111 ANSWER 30 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2005:648160 CAPLUS

143:248607 DN

- Design, Synthesis, and Antiviral Activity of 2'-Deoxy-2'-fluoro-2'-Cmethyl-cytidine, a Potent Inhibitor of Hepatitis C Virus Replication
- Clark, Jeremy L.; Hollecker, Laurent; Mason, J. Christian; Stuyver, Lieven AU J.; Tharnish, Phillip M.; Lostia, Stefania; McBrayer, Tamara R.; Schinazi, Raymond F.; Watanabe, Kyoichi A.; Otto, Michael J.; Furman, Phillip A.; Stec, Wojciech J.; Patterson, Steven E.; Pankiewicz, Krzysztof W. Fharmasset, Inc., Princeton, NJ, 08540, USA Journal of Medicinal Chemistry (2005), 48(17), 5504-5508

- 30
- CODEN: JMCMAR; ISSN: 0022-2623 PB American Chemical Society
- Journal
- T.A English
- os CASREACT 143:248607
- The pyrimidine nucleoside- β-D-2'-deoxy-2'-fluoro-2'-C-methylcytidine
 - (I) was designed as a hepatitis C virus RNA-dependent RNA polymerase (HCV RdRp) inhibitor. The title compound was obtained by a DAST fluorination of N4-benzoy1-1-(2-methy1-3,5-di-0-benzoy1-β-D-arabinofuranosyl)cytosine to provide N4-benzoy1-1-(2-fluoro-2-

methyl-3,5-di-O-benzoyl-β-D-ribofuranosyl)cytosine. The protected

2'-C-methylcytidine was obtained as a byproduct from the DAST fluorination and allowed for the preparation of two biol. active compds. from a common precursor. Compound I and 2'-C-methyloytidine were assayed in a sub-genomic HCV replicon assay system and found to be potent and selective

inhibitors of HCV replication. Compd.I shows increased inhibitory activity in the HCV replicon assay compared to 2'-C-methylcytidine and low cellular toxicity.

863329-62-8P 863329-64-0P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(design, synthesis via fluorination, and antiviral activity of 2'-deoxy-2'-fluoro-2'-C-methyl-cytidine, a potent inhibitor of Hepatitis C virus replication)

863329-62-8 CAPLUS

Benzamide, N-[1-(3,5-di-O-benzoyl-2-C-methyl-β-D-arabinofuranosyl)-1,2-dihydro-2-oxo-4-pyrimidinyl]- (CA INDEX NAME)

863329-64-0 CAPLUS CN Cytidine, N-benzoyl-2'-C-methyl-, 3',5'-dibenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RE.CNT 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 31 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2005:34765 CAPLUS

DN

142:94074 TI Preparation of modified fluorinated (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside analogs as antiviral agents

TN Clark, Jeremy

Pharmasset, Ltd., Barbados PCT Int. Appl., 228 pp. CODEN: PIXXD2 PA

Patent

		glish																
LIN		TENT	NO.															
PI		2005							0113					472			0040	
		W:	AE, CN, GE, LK, NO, TJ, BW, BY, ES, SK,	AG, CO, GH, LR, NZ, TM, GH, KG, FI,	AL, CR, GM, LS, OM, TN, GM, KZ, FR,	AM, CU, HR, LT, PG, TR, KE, MD, GB,	AT, CZ, HU, LU, PH, TT, LS, RU, GR,	AU, DE, ID, LV, PL, TZ, MW, TJ, HU, CG,	AZ, DK, IL, MA, PT, UA, MZ, TM, IE,	BA, DM, IN, MD, RO, UG, SD, AT, IT,	DZ, IS, MG, RU, US, SL, BE, LU,	EC, JP, MK, SC, UZ, SZ, BG, MC,	EE, KE, MN, SD, VC, TZ, CH, NL,	EG, KG, MW, SE, VN, UG, CY, PL,	ES, KP, MX, SG, YU, ZM, CZ, PT,	FI, KR, MZ, SK, ZA, ZW, DE, RO,	GB, KZ, NA, SL, ZM, AM, DK, SE,	GD, LC, NI, SY, ZW, AZ, EE, SI,
	AU	2004				A2		2005	0113		AU 2	004-	2538	60		2	0040	421
	AU	2004	2538	60		A1		2005	0113									
	CA	2527	657			A1		2005	0113		CA 2	004-	2527	657		2	0040	421
	US	2005	0009	737		A1		2005	0113		US 2	004-	8287	53		2	0040	421
	EP	1633	766			A2		2006	0315		EP 2	004-	7759	00		2	0040	421
		R:						ES,										
								RO,										
	BR	2004	0108	46		A		2006	0627		BR 2	004-	1084	6		2	0040	421
	CN	1816 2006	558			A		2006	0809		CN 2	004-	8001	9148		2	0040	421
	JP	2006	5266	29		T		2006	1124		JP 2	006-	5132	31		2	0040	421
	MX	2005	PA12	788		A		2006	0222		MX 2	005-	PA12	788		2	0051	125

IN 2005DN06087 A 20080509 IN 2005-DN6087 20051227

10/609.298

	NO 2005006221	A	20051228	NO	2005-6221	20051228
	US 20080070861	A1	20080320	US	2007-854218	20070912
PRAI	US 2003-474368P	P	20030530			
	US 2004-828753	A3	20040421			
	WO 2004-US12472	W	20040421			
os	MARPAT 142:94074					

The disclosed invention provides nucleoside analogs I, wherein B is purine and pyrimidine nucleobase; X is O, S, CH2, Se, NH, N-alkyl, CHW, C(W)2; W is F, Cl. Br, iodo; R1 is H, phosphate, H-phosphonate, acyl, Ph, alkyl, carboxyalkylamino, sulfonate ester, peptide, amino acid, sugar reside; R2 Carlodyanyananino, surfinite ester, peptue, aminio ant, sugar lester, ac and R2' are independently H, alkyl, alkenyl, alkynyl, vunyl, N3, CN, halogen, NOZ, ester, alkoxy, thioalkyl, sulfoxide, sulfoxyl; R6 is alkyl, CN, Me, OMe, OEt, CH2OH, CH2F, N3, CHCN, CH2NB2, CH2NH2, CH2NHE, CH2NH2, alkylne; and methods of treating a Flavivitidae infection, including hepatitis C virus, West Nile Virus, yellow fever virus, and a rhinovirus infection in a host, including animals, and especially human, using a (2'R)-2'-deoxy-2'-fluoro-2'-C-Me nucleosides, or a pharmaceutically acceptable salt or prodrug thereof. Thus, (2'R)-2'-deoxy-2'-fluoro-2'-C-methylcytidine was prepared and tested as antiviral agent. The effects the nucleoside analogs tested on human bone marrow cells are reported. (2'R)-2'-deoxy-2'-fluoro-2'-C-methylcytidine shows activity against Rhinovirus, West Nile virus, Yellow Fever virus, and Dengue virus. Cytotoxicity and effect of nucleoside analogs on human bone marrow cells are reported.

817204-36-7P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of modified fluorinated (2'R)-2'-deoxy-2'-fluoro-2'-C-Me nucleoside analogs as antiviral agents) 817204-36-7 CAPLUS

Benzamide, N-[1,2-dihydro-1-[2-C-methyl-3,5-bis-0-(trifluoroacetyl)-6-D-arabinofuranosyl]-2-oxo-4-pyrimidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- 111 ANSWER 32 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:780543 CAPLUS
- 141:296247
- Preparation of cytidine nucleoside analogs as antiviral agents
- Girardet, Jean-Luc; Koh, Yung-Hyo; An, Haoyun; Hong, Zhi Ribapharm Inc., USA IN
- οa
- SO PCT Int. Appl., 59 pp.
- Patent
- 1.2 English FAN.CNT 1

	PAI	ENT :	NO.			KIN	0	DATE			APPL	ICAT	ION	NO.		D	ATE	
PI	WO	2004	0804	56		Al		2004	0923		WO 2	003-	US69	92		21	0030	307
		W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
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			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
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		RW:						MZ,										
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					CF,			CM,										
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PRAI		2003				A		2003	0307									
OS GI	MAI	RPAT	141::	2962	47													

- AB Cytidine analogs I, wherein -Xev- is -NeV, -GH-Nr, -NeCZ- or -GH-CZ-, wherein Z is B, halogen, or alkyl, and wherein N is a sugar or sugar analog; wherein the compound has a D-configuration or an L-configuration, with the proviso that where N is a substituted sugar with a ribofurance ring having a heterostom and substituents R1 and R2 on the C3'-atom, R3 and R4 on the R4 o
- IT 23643-36-9 RL: RCI (Reactant); RACI (Reactant or reagent)
- (preparation of cytidine nucleoside analogs as antiviral agents) NN 23643-36-9 CAPLUS
- CN 2,4(1H,3H)-Pyrimidinedione, 1-(2,3,5-tri-O-benzoyl-2-C-methyl-β-D-ribofuranosyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT RE.CNT 9

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L11 ANSWER 33 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN AN 2004\!:\!566635 CAPLUS
DN
        141:89323
       Process for the production of 3'-nucleoside prodrugs
Storer, Richard; Moussa, Adel; Mathieu, Steven; Qu, Lin
IN
        Idenix Cayman Limited, Cayman I.
PCT Int. Appl., 57 pp.
CODEN: PIXXD2
SO
DT
        Patent
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LA English

FAN.		1 TENT	NO.			KIN	D	DATE		APPL	ICAT	ION :	NO.	D	ATE		
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PRAI		2002															
		2003															
os	CA	SREAC	1 14	T:88	323;	MAR	PAT	141:	89323								

GI

Provided is a single-step process for the regioselective 3'-acylation of a ribofuranosyl 2'- or 3'-branched nucleosides I, wherein B is nucleobase.

These compds. are useful as antiviral agents, and in particular, can be used to treat Flaviviridae infections in a host in need thereof (no data). Thus, $9-(2"-C-methyl-3"-O-valinoyl-\beta-D-ribofuranosyl)-6-N$ methyladenine dihydrochloride was prepared via regioselective esterification of 9-(2'-C-methyl-β-D-ribofuranosyl)-6-N-methyladenine with N-(tert-butoxycarbonyl)-L-valine. 640725-70-8P

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(process for production of nucleoside prodrugs via regioselective esterification)

RN

640723-70-8 CAPUS L-Valane, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methyloytidine (CA INDEX NAME)

Absolute stereochemistry.

L11 ANSWER 34 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

AN 2004:453348 CAPLUS 141:17578 DM

Treatment of Flaviviridae infection with 2'-branched nucleosides

and another mutation-inducing drug such as interferon Sommadossi, Jean-Pierre; La Colla, Paolo; Standring, David; Bichko, Vadim; IN

Qu, Lin PA Idenix (Cayman) Limited, Cayman I.; Universita Degli Studi Di Cagliari PCT Int. Appl., 166 pp. CODEN: PIXXD2

DТ Patent

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PRAI		2002						2002	1115										
	WO	2003	-US3	6714		W		2003	1117										

10/609.298

- MARDAT 141 17578
- The present invention discloses a method for the treatment of Flaviviridae infection that includes the administration of a 2'-branched nucleoside, or a pharmaceutically acceptable prodrug and/or salt thereof, to a human in need of therapy in combination or alternation with a drug that directly or indirectly induces a mutation in the viral genome at a location other than a mutation of a nucleotide that results in a change from serine to a different amino acid in the highly conserved consensus sequence, XRX<u>S</u>SGXXXT, of domain B of the RNA polymerase region, or is associated with such a mutation. The invention also includes a method to detect a mutant strain of Flaviviridae and a method for its treatment. Thus, in bovine viral diarrhea virus (BVDV)-infected MDBK cells treated with \$B-D-2'-methylcytidine, viruses resistant to the nucleoside appeared. The drug resistance was associated with a mutation in the NS5B gene which resulted in an \$405T substitution in the encoded RNA-dependent RNA polymerase. These mutant viruses were sensitive to Intron A (interferon α -2b). Intron A and β -D-2'-methylcytidine exhibited synergistic inhibitory activity on BVDV growth in MDBK cells. 640281-90-9
 - RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (treatment of Flaviviridae infection with 2'-branched nucleosides and another mutation-inducing drug such as interferon)
- 640281-90-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

- ANSWER 35 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:20697 CAPLUS 140:87662 DN
- 2'- and 3'-nucleoside prodrugs for treating Flaviviridae infections
- Sommadossi, Jean-pierre; La Colla, Paolo; Storer, Richard; Gosselin, Gilles
- Idenix (Cayman) Limited, Cayman I.; Centre National de la Recherche PA Scientifique; Universita Degli Studi di Cagliari
- PCT Int. Appl., 2498 pp. CODEN: PIXXD2
- Patent

LA FAN.		glish 4																
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     WO 2004-US15395
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os
     MARPAT 140:87662
    2' And 3'-Prodrugs of 1'-, 2'-, 3'-, or 4'-branched \beta-D or \beta-L nucleosides, or their pharmaceutically acceptable salts and derive, are
     described which are useful in the prevention and treatment of
     Flaviviridae infections and other related conditions. These modified nucleosides provide superior results against flaviviruses
     and pestiviruses, including hepatitis C
     virus and viruses generally that replicate through an RNA-dependent RNA
     reverse transcriptase. Compds., compns., methods and uses are provided for the treatment of Flaviviridae infection, including
     HCV infection, that include the administration of an effective
     amount of the prodrugs of the invention, or their pharmaceutically
     acceptable salts or derivs. These drugs may optionally be administered in
     combination or alternation with further antiviral agents to prevent or
     treat Flaviviridae infections and other related conditions.
     Preparation of compds. of the invention is included.
     640725-71-9P
     RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic
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INDEX NAME)

preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(nucleoside prodrugs for treating Flaviviridae infections)

640725-71-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA CN

Absolute stereochemistry. Rotation (+).

●2 HCl

- 640725-69-5P 640725-70-8P
- RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (nucleoside prodrugs for treating Flaviviridae infections)
- 640725-69-5 CAPLUS Cytidine, 2'-C-methyl-, 2',3',5'-tribenzoate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

- 640725-70-8 CAPLUS
- L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

L11 ANSWER 36 OF 37 CAPLUS COPYRIGHT 2008 ACS on STN

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ΔN
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     140:77365
     Preparation of modified 2'- and 3'-nucleoside prodrugs for treating
     Flaviviridae infections
     Sommadossi, Jean-pierre; La Colla, Poalo; Storer, Richard; Gosselin,
    Idenix (Cayman) Limited, Cayman I.; Universita degli studi di Cagliari;
PA
     Centre National de la Recherche Scientifique
SO
    PCT Int. Appl., 201 pp.
     CODEN: PIXXD2
DT
     Patent
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     English
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US 2004-5466

TIS 2004-5472

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US 20070060504

MX 2004PA12709

US 20070060541

TIS 20070060505

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	MARPAT 140:77365					
GI						

2' And/or 3' prodrugs of 1', 2', 3' or 4'-branched-nucleosides I, wherein R1-R3 are independently H, phosphate, alkyl, acyl, C0-alkyl, C0-aryl, C0-alkoxyalkyl, C0-aryloxyalkyl, C0-substituted aryl, sulfonate ester, benzyl, wherein the Ph group is optionally substituted with one or more substituents, alkylsulfonyl, arylsulfonyl, aralkylsulfonyl, lipid, amino acid, carbohydrate, peptide, cholesterol; Y1 is hydrogen, bromo, chloro, fluoro, iodo, CN, OH, OR4, NH2, NH84, NR485, SH or SR4; X1 and X2 are independently alkyl, CH3, CF3, C73, 2-B-TET, CREF, CH2Cl, CH2CF3, CF2CF3, CY2CY3, CH2OH, alkenyl, alkynyl, COOH, COOR4, COO-alkyl, COO-aryl, COO-alkoxyalkyl, CONH2, CONH24, CON(34)2, halo, CN, N3, OH, OR4, NH2, NH4, NR45, SH or SR5; Y is independently H, halo; and each R4 and R5 is name, names, on to say, i is independently n, hady and weather and a lindependently hydrogen, acyl, alkyl, lower alkyl, alkenyl, alkynyl or cycloalkyl, and their pharmaceutically acceptable salts and derive. are described. These prodrugs are useful in the prevention and treatment of Flaviviriade infections, including HCV infection, and other related conditions. Compds. and compns. of the prodrugs of the present invention are described. Methods and uses are also provided that include the administration of an effective amount of the prodrugs of the present invention, or their pharmaceutically acceptable salts or derivs. These drugs may optionally be administered in combination or alteration with further anti-viral agents to prevent or treat Flaviviridae infections and other related conditions. Thus, antiviral activity of β-D-2'-C-methyl-7-methyl-6-phenyl-3,3a,5,8a-tetrahydro-1,3,4,5,7apenta-aza-s-indacen-8-one is reported. 640281-90-9P

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of nodified and nucleoside prodrugs for treating flaviviridae infections) RN 64028-9-9-0 (2011);

RN 640281-90-9 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Lil ANSWER 37 OF 37 CAPLUS COFYRIGHT 2008 ACS on STN
AN 2004:20443 CAPLUS
DN 140:79941-3'-0-L-valine ester ribofuranceyl cytidine for treatment of
filaviviticae infections
IN Sommadorsi, Jean-Pierre; la Colla, Paolo
Fa Idenix (Cayman) Linted, Cayman I.; Universita Degli Studi di Cagliari
FO PT Int. Appl., 110 pp.
CODEN: PIERCS
CODEN: PIERCS

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     IN 2005-DN344
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     MARPAT 140:70984
     The 3'-L-valine ester of β-D-2'-C-methyl-ribofuranosyl cytidine
     provides superior results against flaviviruses and
     pestiviruses, including hepatitis C virus.
     Based on this discovery, compds., compns., methods and uses are provided
     for the treatment of flaviviridae, including HCV, that
     include the administration of an effective amount of val-mCyd or its salt,
     ester, prodrug or derivative, optionally in a pharmaceutically acceptable
     carrier. In an alternative embodiment, val-mCyd is used to treat any
     virus that replicates through an RNA-dependent RNA polymerase. Several
     examples are provided of the pharmacol., mechanism of action, metabolism, side effects, and clin. efficacy of the title compound
     640281-90-9D, salts 642075-50-1 642075-51-2
     642075-52-3 642075-53-4 642075-54-5
     642075-55-6 642075-56-7 642075-57-8
     642075-58-9 642075-59-0 642075-60-3
     642075-61-4 642075-62-5 642075-63-6
     642075-64-7 642075-65-8 642075-66-9
     642075-67-0 642075-68-1 642075-69-2
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642075-70-5 642075-71-6 642075-72-7 642075-74-9 642075-75-0 642075-76-1 642075-77-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ribofuranosylcytidine methylvaline ester combined with other antivirals for treatment of flaviviridae infections) 640281-90-9 CAPLUS RN

L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN

642075-50-1 CAPLUS 1-Valine, 3'-ester with 2'-C-methylcytidine, 4-methylbenzenesulfonate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 104-15-4 CMF C7 H8 03 S

RN

642075-51-2 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, methanesulfonate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 75-75-2 CMF C H4 03 S

642075-52-3 CAPLUS 1-Valine, 3'-ester with 2'-C-methylcytidine, acetate (salt) (9CI) (CA

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10/609,298
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INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 64-19-7 CMF C2 H4 02

642075-53-4 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxy-1,2,3-propanetricarboxylate (salt) (9CI) (CA INDEX NAME)

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM

CRN 77-92-9 CMF C6 H8 07

CO2H

но2с-сн2-с-сн2-со2н OH

642075-54-5 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, propanedioate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM

CRN 141-82-2 CMF C3 H4 O4

HO2C-CH2-CO2H

642075-55-6 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, (2R, 3R)-2, 3-dihydroxphutanedioate (salt) (9CI) (CA INDEX NAME) CN

CM

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 87-69-4 CMF C4 H6 O6

Absolute stereochemistry.

642075-56-7 CAPLUS L-Valline, 3'-ester with 2'-C-methylcytidine, butanedioate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM

CRN 110-15-6 CMF C4 H6 O4

HO2C-CH2-CH2-CO2H

RN 642075-57-8 CAPLUS L-Valine, 3'-ester with 2'-C-methyloytidine, benzoate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 65-85-0 CMF C7 H6 O2

642075-58-9 CAPLUS L-Ascorbic acid, compd. with L-valine 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 50-81-7 CMF C6 H8 O6

Absolute stereochemistry.

RN

642075-59-0 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, 2-oxopentanedioate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM

CRN 328-50-7 CMF C5 H6 O5

642075-60-3 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, 2,3-dihydroxypropyl phosphate (salt) (9CI) (CA INDEX NAME)

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10/609,298

CM 1

CRN 640281-90-9

CMF CL5 H24 N4 06

Absolute Stereochemistry. Rotation (+).
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CRN 57-03-4 CMF C3 H9 O6 P

οн

Ho-CH2-CH-CH2-OPO3H2

RN 642075-61-4 CAPLUS

ND L-Valine, 3'-ester with 2'-C-methyloytidine, formate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 64-18-6 CMF C H2 O2

о== сн- он

RN 642075-62-5 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methyloytidine, (2E)-2-butenedioate (salt)
(961) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 O6 Absolute stereochemistry. Rotation (+).

CM

CRN 110-17-8 CMF C4 H4 04

Double bond geometry as shown.

642075-63-6 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine, propanoate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9

CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 79-09-4 CMF C3 H6 02

но-с-сн2-сн3

642075-64-7 CAPLUS

L-Valine, 3'-ester with 2'-C-methyloytidine, hydroxyacetate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

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CM
      CRN 79-14-1
      CMF C2 H4 03
но-с-сн2-он
    642075-65-8 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxypropanoate (salt) (9C1) (CA INDEX NAME)
      CM 1
      CRN 640281-90-9
CMF C15 H24 N4 06
Absolute stereochemistry. Rotation (+).
      CM 2
      CRN 50-21-5
      CMF C3 H6 O3
    OH
Me-CH-CO2H
RN
    642075-66-9 CAPLUS
1-Valine, 3'-ester with 2'-C-methylcytidine, 2-oxopropanoate (salt) (9CI)
(CA INDEX NAME)
CN
      CM 1
      CRN 640281-90-9
CMF C15 H24 N4 06
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McIntosh

McIntosh

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

642075-69-2 CAPLUS L-Valine, 3'-ecter with 2'-C-methylcytidine, 2-hydroxybenzoate (salt) (9C1) (CA INDEX NAME)

CM

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 69-72-7 CMF C7 H6 03

642075-70-5 CAPLUS L-Valine, 3'-ester with 2'-C-methyloytidine, sulfate (salt) (9CI) (CA INDEX NAME) CN

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

CRN 7664-93-9 CMF H2 04 S

но- s- он

642075-71-6 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, nitrate (salt) (9CI) (CA INDEX NAME)

CM

CRN 640281-90-9

CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

> CRN 7697-37-2 CMF H N 03

642075-72-7 CAPLUS

L-Valine, 3'-ester with 2'-C-methyloytidine, carbonate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

CRN 463-79-6 CMF C H2 03

642075-74-9 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydrobromide (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

•x HBr

642075-75-0 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, hydriodide (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

•x HI

642075-76-1 CAPLUS L-Valine, 3'-ester with 2'-C-methylcytidine, carbonate (2:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 O6

Absolute stereochemistry. Rotation (+).

CM 2

CRN 463-79-6 CMF C H2 O3

HO-C-OH

RN 642075-77-2 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, phosphate (salt) (9CI) (CA
INDEX NAME)

CM 1

CRN 640281-90-9 CMF C15 H24 N4 06

Absolute stereochemistry. Rotation (+).

CM 2

CRN 7664-38-2 CMF H3 04 P

0 | |-|-

OH T 640281-90-9P

RL: DNA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (ribofuranceylcytidine methylvaline ester for treatment of flaviviriade infections)

640281-90-9 CAPLUS

L-Valine, 3'-ester with 2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

642075-41-0

RL: RCT (Reactant); RACT (Reactant or reagent)

(ribofuranosylcytidine methylvaline ester for treatment of flaviviridae infections)

RN

642075-41-0 CAPLUS Cytidine, 2'-C-methyl-, 2',3',5'-tris(benzeneacetate) (9CI) (CA INDEX NAMEL

Absolute stereochemistry.

640725-70-8P 642075-44-3P 642075-48-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(ribofurancsylcytidine methylvaline ester for treatment of flaviviridae infections)

640725-70-8 CAPLUS L-Valine, N-[(1,1-dimethylethoxy)carbonyl]-, 3'-ester with

2'-C-methylcytidine (CA INDEX NAME)

Absolute stereochemistry.

642075-44-3 CAPLUS

Owner of the second sec

Absolute stereochemistry. Double bond geometry unknown.

642075-48-7 CAPLUS Cytidine, 2'-C-methyl-N-(phenylacetyl)-, 2',3',5'-tris(benzeneacetate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

640725-71-9P

Gaulia-12-32
RI: SPN (Gynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREF (Preparation); USES (Uses) (ribofurancsylcytidin methylvaline ester for treatment of flaviviridae infections)

RN

riaviviriame infections)
640725-71-9 CAPIUS
L-Valine, 3'-ester with 2'-C-methylcytidine, hydrochloride (1:2) (CA
INDEX NAME)

Absolute stereochemistry. Rotation (+).

● 2 HCl

	(FILE	E 'HOME' ENTERED AT 18:07:49 ON 08 JUN 2008)
11 12 13	FILE	'REGISTRY' ENTERED AT 18:07:59 ON 08 JUN 2008 STRUCTURE UPLOADED 24 S LI 519 S LI FULL
14 15 16	FILE	'CAPLUS' ENTERED AT 18:08:46 ON 08 JUN 2008 120 5 L3 22498 5 L4 AND FLAVIVIRUS OR PESTIVIRUS OR FLAVIVIRIDAE OR HCV OR HEP 58 5 L4 AND (FLAVIVIRUS OR PESTIVIRUS OR FLAVIVIRIDAE OR HCV OR HE
17 18 19	FILE	'REGISIRY' ENTERED AT 18:13:47 ON 08 JUN 2008 STRUCTURE UPLOADED 4 S L7 171 S L7 FULL
L10 L11	FILE	'CAPLUS' ENTERED AI 18:14:23 ON 08 JUN 2008 74 S L9 37 S L10 AND (FLAVIVIRUS OR PESTIVIRUS OR FLAVIVIRIDAE OR HCV OR H